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(54) Simultaneous detection, identification and differentiation of Eubacterial taxa using a hybridization assay

(57) The present invention relates to a method for detection and identification of at least one micro-organism, or for the simultaneous detection of several micro-organisms in a sample, comprising the steps of:

- (i) if need be releasing, isolating or concentrating the polynucleic acids present in the sample;
- (ii) if need be amplifying the 16S-23S rRNA spacer region, or a part of it, with at least one suitable primer pair;
- (iii) hybridizing the polynucleic acids of step (i) or (ii) with at least one and preferably more than one of the spacer probes as mentioned in table Ia or equivalents of thereof, under the appropriate hybridization

and wash conditions, and/or with a taxon-specific probe derived from any of the spacer sequences as represented in figs. 1-103 under the same hybridization and wash conditions;
(iv) detecting the hybrids formed in step (iii) with each of the probes used under appropriate hybridization and wash conditions;
(v) identification of the micro-organism(s) present in the sample from the differential hybridization signals obtained in step (iv).

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Description

[0001] The present invention relates to nucleic acid probes derived from the spacer region between the 16S and 23S ribosomal ribonucleic acid (rRNA) genes, to be used for the specific detection of eubacterial organisms in a biological sample by a hybridization procedure, as well as to nucleic acid primers to be used for the amplification of said spacer region of eubacterial organisms in a biological sample. The present invention also relates to new spacer region sequences from which said probes or primers may be derived.

[0002] Since the advent of the polymerase chain reaction and some other nucleic acid amplification techniques the impact of DNA-probe technology in the diagnosis of micro-organisms in biological samples of all sorts is increasing. Being often more specific and potentially more sensitive - if an adequate amplification and/or detection system is used - the DNA probe approach may eventually replace the conventional identification techniques.

[0003] The reliability of nucleic acid based tests essentially depends on the sensitivity and specificity of the probes and/or primers used. Thus the corner stone of this type of assay is the identification of nucleic acid sequences which are unique to the group of organisms of interest.

[0004] Most of the nucleic acid based tests either described in literature and/or commercially available aim at the detection of just one particular organism in a biological sample. Since most biological samples usually may contain a great variety of clinically relevant micro-organisms, a multitude of separate assays have to be performed to detect all relevant organisms possibly present. This approach would be very expensive, laborious and time-consuming. Consequently, the number of tests actually performed in most routine diagnostic labs on a particular sample is restricted to the detection of just a few of the most relevant organisms. Therefore it would be extremely convenient to have access to a system which enables the fast, easy and simultaneous detection of a multitude of different organisms. The more organisms that can be screened for in the same assay, the more cost-effective the procedure would be.

[0005] As put forward in earlier published documents, the spacer region situated between the 16S rRNA and the 23S rRNA gene, also referred to as the internal transcribed spacer (ITS), is an advantageous target region for probe development for detection of pathogens of bacterial origin (International application WO 91/16454; Rossau et al., 1992; EP-A-0 395 292).

[0006] One of its most appreciated advantages is that sequences unique to a great variety of bacterial taxa can be found in a very limited area of the bacterial genome. This characteristic allows for an advantageous design of "probe-pans" enabling the simultaneous detection of a set of organisms possibly present in a particular type of a biological sample. Moreover, being flanked by quasi-universally conserved nucleotide sequences - more particularly located in the 3'-part of the 16S rRNA gene and the 5'-part of the 23S rRNA gene respectively - almost all spacers can be simultaneously amplified with a limited set of amplification primers. Alternatively, specific primer sets can be derived from the spacer sequences themselves, thereby allowing species- or group-specific amplifications.

[0007] The 16S-23S rRNA spacer region is a relatively short (about 200 to 1000 base pairs) stretch of DNA present in one or multiple copies in the genome of almost all eubacterial organisms. If multiple copies are present in the genome of one bacterium these copies can either be identical (as is most probably the case in some Neisseria species) or may differ from each other (as is the case for E. coli). This difference can be limited to a few nucleotides but also deletions and insertions of considerable length may be present.

[0008] Until now, spacer probes are only described and made publicly available for a limited number of organisms many of which were disclosed in international application WO 91/16454. As described above, it would be very advantageous to be able to detect simultaneously a panel of pathogens: e.g. a panel of pathogens possibly present in the same type of biological sample, or a panel of pathogens possibly causing the same type of disease symptoms, which are difficult to differentiate clinically and/or biochemically, or a panel of organisms belonging to the same taxon. In order to make the different panels as complete as possible, additional probes or sets of probes located in the spacer region and enabling the identification of at least the following bacterial groups or species are required:

- Mycobacterium species
- Listeria species
- Chlamydia species
- Acinetobacter species
- Mycoplasma species
- Streptococcus species
- Staphylococcus species
- Salmonella species
- Brucella species
- Yersinia species
- Pseudomonas species

[0009] These additional spacer probes need to be meticulously designed such that they can be used simultaneously with at least one other probe, under the same hybridization and wash conditions, allowing the detection of a particular panel of organisms.

[0010] It is thus the aim of the present invention to select probes or sets of probes, which have as target the 16S-23S rRNA spacer region, and which allow the detection and identification of at least one, and preferably more than one, of the above mentioned micro-organisms. The probes or probe sets are selected in such a way that they can be used in combination with at least one other probe, preferably also originating from the 16S-23S rRNA spacer region, under the same hybridisation and wash conditions, to allow possibly the simultaneous detection of several micro-organisms in a sample.

[0011] It is also an aim of the present invention to provide for a selection method for use in the selection of said spacer probes or probe sets.

[0012] It is also an aim of the present invention to provide a rapid and reliable hybridization method for detection and identification of at least one micro-organism in a sample, or for the simultaneous detection and identification of several micro-organisms in a sample.

[0013] It is more particularly an aim of the present invention to provide a hybridization method allowing simultaneous detection and identification of a set of micro-organisms, liable to be present in a particular type of sample.

[0014] It is more particularly an aim of the present invention to provide probes or sets of probes for the possible simultaneous detection of micro-organisms in a sample originating from respiratory tract.

[0015] It is another particular aim of the present invention to provide probes or sets of probes for the possible simultaneous detection of micro-organisms in a sample originating from cerebrospinal fluid.

[0016] It is still another particular aim of the present invention to provide probes or sets of probes for the possible simultaneous detection of micro-organisms in a sample originating from urogenital tract.

[0017] It is still another particular aim of the present invention to provide probes or sets of probes for the possible simultaneous detection of micro-organisms in a sample taken from the gastro-intestinal tract of a patient.

[0018] It is still another particular aim of the present invention to provide probes or sets of probes for the possible simultaneous detection of micro-organisms in a sample originating from food or environmental samples.

[0019] It is moreover an aim of the present invention to provide a method for detection and identification of a particular taxon in a sample, or a set of particular taxa, said taxon being either a complete genus, or a subgroup within a genus, a species or even subtypes within a species (subspecies, serovars, sequevars, biovars...).

[0020] It is more particularly an aim of the present invention to provide probes or sets of probes for the detection of Mycobacterium species and subspecies, more particularly for the detection of M. tuberculosis complex strains, Mycobacterium strains from the MAIS-complex, M. avium and M. paratuberculosis, M. intracellulare and M. intracellulare-like strains, M. scrofulaceum, M. kansasii, M. chelonae, M. gordonae, M. ulcerans, M. genavense, M. xenopi, M. simiae, M. fortuitum, M. malmoense, M. celatum and M. haemophilum.

[0021] It is also an aim of the present invention to provide probes or sets of probes for the detection of Mycoplasma strains, more particularly of M. pneumoniae and M. genitalium.

[0022] It is also an aim of the present invention to provide probes or sets of probes for the detection of Pseudomonas strains, more particularly P. aeruginosa.

[0023] It is also an aim of the present invention to provide probes or sets of probes for detection of Staphylococcus species, more particularly S. aureus and S. epidermidis.

[0024] It is also an aim of the present invention to provide probes or sets of probes for the detection of Acinetobacter strains, more particularly A. baumanii.

[0025] It is also an aim of the present invention to provide probes or sets of probes for the detection of Listeria strains, more particularly Listeria monocytogenes.

[0026] It is also an aim of the present invention to provide probes or sets of probes for the detection of Brucella strains.

[0027] It is also an aim of the present invention to provide probes or sets of probes for the detection of Salmonella strains.

[0028] It is also an aim of the present invention to provide probes or sets of probes for the detection of Chlamydia strains, more particularly C. trachomatis and C. psittaci.

[0029] It is also an aim of the present invention to provide probes or sets of probes for the detection of Streptococcus strains.

[0030] It is also an aim of the present invention to provide probes or sets of probes for the detection of Yersinia enterolitica strains.

[0031] It is also an aim of the present invention to provide primers allowing specific amplification of the 16S-23S rRNA spacer region for certain organisms. More particularly, it is an aim of the present invention to provide primers for the specific amplification of the spacer region of Mycobacterium, Chlamydia, Listeria, Brucella and Yersinia enterolitica strains.

[0032] It is also an aim of the present invention to provide new sequences of 16S-23S rRNA spacer regions from

which useful spacer probes or primers can be derived.

[0033] It is also an aim of the present invention to provide for kits for detection of at least one organism in a sample in which said probes and/or primers are used.

[0034] It is noted that for a few of the above-mentioned organisms spacer sequences have already been published in literature or in publicly accessible data-banks.

[0035] However, it should be made clear that the spacer region sequences disclosed in the current invention (figs. 1-103) are new and, in case they originate from the same species as those of which a spacer sequence was already described in the prior art, they differ to some extent from the already described sequences.

[0036] Moreover, it is the principal aim of the present invention to select, from the compilation of sequence data on spacer regions, specific probes and sets of probes enabling the detection and identification of a particular panel of organisms, be it the organisms belonging to a common taxon, or the organisms possibly present in the same type of sample.

[0037] The selection procedure usually consists of a theoretical and an experimental part. First of all, the different spacer sequences need to be aligned to those of the 'closest neighbours' or to the spacer sequences of other micro-organisms liable to be present in the same sample. This requires of course the sequence determination of the spacer region, as described in the examples. From the alignment, regions of divergence can be defined, from which probes with desired hybridization characteristics are designed, according to guidelines known to the man skilled in the art and specified in more detail below.

[0038] Secondly, the designed probes need to be tested experimentally and evaluated for their usefulness under specific hybridization conditions and/or in combination with other probes. Experimental testing can be done according to any hybridization method known in the art, but a preferred assay for the simultaneous testing of different probes under the same conditions is the reverse hybridization assay. A specific format for reverse hybridization of different probes simultaneously used in the current invention is the LiPA (Line Probe Assay) as described below.

[0039] Upon experimental testing unexpected hybridization behaviour may show up when the probes are hybridized to the target nucleic acid, and specific probe adaptations may be required.

[0040] Moreover, specificity and sensitivity of the probes need to be tested with a large collection of strains, both belonging to the taxon to be detected and belonging to other taxa. Due to genome heterogeneity in the spacer region, or the existence of multiple spacer regions with different sequences in the same organism, it is quite often necessary to sequence spacer regions of additional strains, or to sequence additional spacer regions in the same strain, and redesign the probes according to the new sequence data in order to obtain a better sensitivity and/or specificity (see e.g. example 3). In some cases it may be necessary or preferable to use several probes for the same organism (see e.g. example 2 and 7). Also, upon sequencing the spacer region, some organisms may show unexpected (un)relatedness, which may lead to a revision of strain classification contrary to classical taxonomic criteria (see e.g. examples 2 and 7).

[0041] In conclusion, the experimental part of the probe selection procedure is indispensable and complementary to the theoretical part. Probe design, especially under the fixed conditions of reverse hybridization (the same conditions for each probe) is not straightforward and probes have to be evaluated meticulously before they can be used in a reverse hybridization format. Therefor, probes cannot always be simply derived on a theoretical basis from a known gene sequence.

[0042] For designing probes with desired characteristics the following useful guidelines may be followed.

[0043] Because the extent and specificity of hybridization reactions such as those described herein are affected by a number of factors, manipulation of one or more of those factors will determine the exact sensitivity and specificity of a particular probe, whether perfectly complementary to its target or not. The importance and effect of various assay conditions, explained further herein, are known to those skilled in the art.

[0044] First, the stability of the [probe : target] nucleic acid hybrid should be chosen to be compatible with the assay conditions. This may be accomplished by avoiding long A and T rich sequences, by terminating the hybrids with G:C base pairs, and by designing the probe with an appropriate Tm. The beginning and end points of the probe should be chosen so that the length and %GC result in a Tm about 2-10°C higher than the temperature at which the final assay will be performed. The base composition of the probe is significant because G-C base pairs exhibit greater thermal stability as compared to A-T base pairs due to additional hydrogen bonding. Thus, hybridization involving complementary nucleic acids of higher G-C content will be stable at higher temperatures.

[0045] Conditions such as ionic strength and incubation temperature under which a probe will be used should also be taken into account in constructing a probe. It is known that hybridization will increase as the ionic strength of the reaction mixture increases, and that the thermal stability of the hybrids will increase with increasing ionic strength. On the other hand, chemical reagents, such as formamide, urea, DMSO and alcohols, which disrupt hydrogen bonds, will increase the stringency of hybridization. Destabilization of the hydrogen bonds by such reagents can greatly reduce the Tm. In general, optimal hybridization for synthetic oligonucleotide probes of about 10-50 bases in length occurs approximately 5°C below the melting temperature for a given duplex. Incubation at temperatures below the optimum

may allow mismatched base sequences to hybridize and can therefore result in reduced specificity.

[0046] It is desirable to have probes which hybridize only under conditions of high stringency. Under high stringency conditions only highly complementary nucleic acid hybrids will form; hybrids without a sufficient degree of complementarity will not form. Accordingly, the stringency of the assay conditions determines the amount of complementarity needed between two nucleic acid strands forming a hybrid. Stringency is chosen to maximize the difference in stability between the hybrid formed with the target and the nontarget nucleic acid. In some examples of the current invention, e.g. when highly related organisms need to be differentiated, it may be necessary to detect single base pair changes. In those cases, conditions of very high stringency are needed.

[0047] Second, probes should be positioned so as to minimize the stability of the [probe : nontarget] nucleic acid hybrid. This may be accomplished by minimizing the length of perfect complementarity to non-target organisms, avoiding GC rich regions of homology to non-target sequences, and by positioning the probe to span as many destabilizing mismatches as possible. Whether a probe sequence is useful to detect only a specific type of organism depends largely on the thermal stability difference between (probe:target) hybrids and [probe:nontarget] hybrids. In designing probes, the differences in these Tm values should be as large as possible (e.g. at least 2°C and preferably 5°C).

[0048] The length of the target nucleic acid sequence and, accordingly, the length of the probe sequence can also be important. In some cases, there may be several sequences from a particular region, varying in location and length, which will yield probes with the desired hybridization characteristics. In other cases, one sequence may be significantly better than another which differs merely by a single base. While it is possible for nucleic acids that are not perfectly complementary to hybridize, the longest stretch of perfectly complementary base sequence will normally primarily determine hybrid stability. While oligonucleotide probes of different lengths and base composition may be used, oligonucleotide probes preferred in this invention are between about 10 to 50 bases in length and are sufficiently homologous to the target nucleic acid.

[0049] Third, regions in the target DNA or RNA which are known to form strong internal structures inhibitory to hybridization are less preferred. Likewise, probes with extensive self-complementarity should be avoided. As explained above, hybridization is the association of two single strands of complementary nucleic acids to form a hydrogen bonded double strand. It is implicit that if one of the two strands is wholly or partially involved in a hybrid that it will be less able to participate in formation of a new hybrid. There can be intramolecular and intermolecular hybrids formed within the molecules of one type of probe if there is sufficient self complementarity. Such structures can be avoided through careful probe design. By designing a probe so that a substantial portion of the sequence of interest is single stranded, the rate and extent of hybridization may be greatly increased. Computer programs are available to search for this type of interaction. However, in certain instances, it may not be possible to avoid this type of interaction.

[0050] The probes of the present invention are designed for attaining optimal performance under the same hybridization conditions so that they can be used in sets for simultaneous hybridization; this highly increases the usability of these probes and results in a significant gain in time and labour. Evidently, when other hybridization conditions should be preferred, all probes should be adapted accordingly by adding or deleting a number of nucleotides at their extremities. It should be understood that these concomitant adaptations should give rise to essentially the same result, namely that the respective probes still hybridize specifically with the defined target. Such adaptations might also be necessary if the amplified material should be RNA in nature and not DNA as in the case for the NASBA system.

[0051] The hybridization conditions can be monitored relying upon several parameters, such as the nature and concentration of the components of the media, and the temperatures under which the hybrids are formed and washed.

[0052] The hybridization and wash temperature is limited in upper value depending on the sequence of the probe (its nucleic acid composition, kind and length). The maximum hybridization or wash temperature of the probes described in the present invention ranges from 40°C to 60°C, more preferably from 45°C to 55°C, in the specific hybridization and wash media as described in the Examples section. At higher temperatures duplexing (= formation of the hybrids) competes with the dissociation (or denaturation) of the hybrid formed between the probe and the target.

[0053] In a preferred hybridization medium of the invention, containing 3 x SSC and 20% formamide, hybridization temperatures can range from 45°C to 55°C, with a preferred hybridization temperature of 50°C. A preferred wash medium contains 3 x SSC and 20% formamide, and preferred wash temperatures are the same as the preferred hybridization temperatures, i.e. preferably between 45°C and 55°C, and most preferably 50°C.

[0054] However, when modifications are introduced, be it either in the probes or in the media, the temperatures at which the probes can be used to obtain the required specificity should be changed according to known relationships, such as those described in the following reference: Hames B and Higgins S (eds.). Nucleic acid hybridization. A practical approach, IRL Press, Oxford, U.K., 1985.

[0055] The selected nucleic acid probes derived from the 16S-23S rRNA spacer region and described by the present invention are listed in Table Ia (SEQ ID NO 1 to 64, 175 to 191, 193 to 201, and 210 to 212). As described in the examples section, some of these probes show a better sensitivity and/or specificity than others, and the better probes are therefore preferentially used in methods to detect the organism of interest in a biological sample. However, it is possible that for certain applications (e.g. epidemiology, substrate typing, ...) a set of probes including the less specific

and/or less sensitive probes may be very informative (see e.g. example 7).

[0056] The following definitions serve to illustrate the terms and expressions used in the different embodiments of the present invention as set out below.

5 [0057] The term "spacer" is an abbreviated term referring to the 16S-23S rRNA internal transcribed spacer region.

[0058] The term "probe" refers to single stranded sequence-specific oligonucleotides which have a sequence which is sufficiently complementary to hybridize to the target sequence to be detected.

[0059] The more specific term "spacer probe" refers to a probe as defined above having a sequence which is sufficiently complementary to hybridize to a target sequence which is located in the spacer region(s) of the organism (or group of organisms) to be detected.

10 [0060] Preferably said probes are 70%, 80%, 90%, or more than 95% homologous to the exact complement of the target sequence to be detected. These target sequences are either genomic DNA or precursor RNA, or amplified versions thereof.

15 [0061] Preferably, these probes are about 5 to 50 nucleotides long, more preferably from about 10 to 25 nucleotides. The nucleotides as used in the present invention may be ribonucleotides, deoxyribonucleotides and modified nucleotides such as inosine or nucleotides containing modified groups which do not essentially alter their hybridization characteristics. Moreover, it is obvious to the man skilled in the art that any of the below-specified probes can be used as such, or in their complementary form, or in their RNA form (wherein T is replaced by U).

20 [0062] The probes according to the invention can be formed by cloning of recombinant plasmids containing inserts including the corresponding nucleotide sequences, if need be by cleaving the latter out from the cloned plasmids upon using the adequate nucleases and recovering them, e.g. by fractionation according to molecular weight. The probes according to the present invention can also be synthesized chemically, for instance by the conventional phospho-ester method.

25 [0063] The term "complementary" nucleic acids as used herein means that the nucleic acid sequences can form a perfect base-paired double helix with each other.

[0064] The term "homologous" as used in the current application is synonymous for identical: this means that poly-nucleic acids which are said to be e.g. 80% homologous show 80% identical base pairs in the same position upon alignment of the sequences.

30 [0065] The term "polynucleic acid" corresponds to either double-stranded or single-stranded cDNA or genomic DNA or RNA, containing at least 10, 20, 30, 40 or 50 contiguous nucleotides. A polynucleic acid which is smaller than 100 nucleotides in length is often also referred to as an oligonucleotide. Single stranded polynucleic acid sequences are always represented in the current invention from the 5' end to the 3' end.

[0066] The term 'closest neighbour' means the taxon which is known or expected to be most closely related in terms of DNA homology and which has to be differentiated from the organism of interest.

35 [0067] The expression 'desired hybridization characteristics' means that the probe only hybridizes to the DNA or RNA from organisms for which it was designed, and not to DNA or RNA from other organisms (closest neighbours or organisms liable to be present in the same sample). In practice, this means that the intensity of the hybridization signal is at least two, three, four, five, ten or more times stronger with the target DNA or RNA from the organisms for which the probes were designed, as compared to non-target sequences.

[0068] These desired hybridization characteristics correspond to what is called later in the text "specific hybridization".

40 [0069] The expression "taxon-specific hybridization" or "taxon-specific probe" means that the probe only hybridizes to the DNA or RNA from the taxon for which it was designed and not to DNA or RNA from other taxa.

[0070] The term taxon can refer to a complete genus or a sub-group within a genus, a species or even subtype within a species (subspecies, serovars, sequevars, biovars...).

45 [0071] The term "specific amplification" or "specific primers" refers to the fact that said primers only amplify the spacer region from these organisms for which they were designed, and not from other organisms.

[0072] The term "sensitivity" refers to the number of false negatives: i.e. if 1 of the 100 strains to be detected is missed out, the test shows a sensitivity of (100-1/100)% = 99%.

[0073] The term "specificity" refers to the number of false positives: i.e. if on 100 strains detected, 2 seem to belong to organisms for which the test is not designed, the specificity of the test is (100-2/100)% = 98%.

50 [0074] The probes selected as being "preferential" show a sensitivity and specificity of more than 80%, preferably more than 90% and most preferably more than 95%.

[0075] The term "primer" refers to a single stranded DNA oligonucleotide sequence capable of acting as a point of initiation for synthesis of a primer extension product which is complementary to the nucleic acid strand to be copied. The length and the sequence of the primer must be such that they allow to prime the synthesis of the extension products. Preferably the primer is about 5-50 nucleotides long. Specific length and sequence will depend on the complexity of the required DNA or RNA targets, as well as on the conditions of primer use such as temperature and ionic strength. The fact that amplification primers do not have to match exactly with the corresponding template sequence to warrant proper amplification is amply documented in the literature (Kwok et al., 1990).

[0076] The amplification method used can be either polymerase chain reaction (PCR; Saiki et al., 1988), ligase chain reaction (LCR; Landgren et al., 1988; Wu & Wallace, 1989; Barany, 1991), nucleic acid sequence-based amplification (NASBA; Guatelli et al., 1990; Compton, 1991), transcription-based amplification system (TAS; Kwok et al., 1989), strand displacement amplification (SDA; Duck, 1990; Walker et al., 1992) or amplification by means of QB replicase (Lizardi et al., 1988; Lomeli et al., 1989) or any other suitable method to amplify nucleic acid molecules known in the art.

[0077] The oligonucleotides used as primers or probes may also comprise nucleotide analogues such as phosphorothioates (Matsukura et al., 1987), alkylphosphorothioates (Miller et al., 1979) or peptide nucleic acids (Nielsen et al., 1991; Nielsen et al., 1993) or may contain intercalating agents (Asseline et al., 1984).

[0078] As most other variations or modifications introduced into the original DNA sequences of the invention these variations will necessitate adaptions with respect to the conditions under which the oligonucleotide should be used to obtain the required specificity and sensitivity. However the eventual results of hybridisation will be essentially the same as those obtained with the unmodified oligonucleotides.

[0079] The introduction of these modifications may be advantageous in order to positively influence characteristics such as hybridization kinetics, reversibility of the hybrid-formation, biological stability of the oligonucleotide molecules, etc.

[0080] The term "solid support" can refer to any substrate to which an oligonucleotide probe can be coupled, provided that it retains its hybridization characteristics and provided that the background level of hybridization remains low. Usually the solid substrate will be a microtiter plate, a membrane (e.g. nylon or nitrocellulose) or a microsphere (bead). Prior to application to the membrane or fixation it may be convenient to modify the nucleic acid probe in order to facilitate fixation or improve the hybridization efficiency. Such modifications may encompass homopolymer tailing, coupling with different reactive groups such as aliphatic groups, NH₂ groups, SH groups, carboxylic groups, or coupling with biotin, haptens or proteins.

[0081] The term "labelled" refers to the use of labelled nucleic acids. Labelling may be carried out by the use of labelled nucleotides incorporated during the polymerase step of the amplification such as illustrated by Saiki et al. (1988) or Bej et al. (1990) or by the use of labelled primers, or by any other method known to the person skilled in the art. The nature of the label may be isotopic (³²P, ³⁵S, etc.) or non-isotopic (biotin, digoxigenin, etc.).

[0082] The "sample" may be any biological material taken either directly from the infected human being (or animal), or after culturing (enrichment), or a sample taken from food or feed. Biological material may be e.g. expectorations of any kind, bronchoolavages, blood, skin tissue, biopsies, lymphocyte blood culture material, colonies, etc. Said samples may be prepared or extracted according to any of the techniques known in the art.

[0083] The "target" material in these samples may be either genomic DNA or precursor RNA of the organism to be detected (=target organism), or amplified versions thereof as set out above. More specifically, the nucleic acid sequence of the target material is localized in the spacer region of the target organism(s).

[0084] Detection and identification of the target material can be performed by using one of the many electrophoresis methods, hybridization methods or sequencing methods described in literature and currently known by men skilled in the art. However, a very convenient and advantageous technique for the simultaneous detection of nucleic acids possibly present in biological samples is the Line Probe Assay technique. The Line Probe Assay (LiPA) is a reverse hybridization format (Saiki et al., 1989) using membrane strips onto which several oligonucleotide probes (including negative or positive control oligonucleotides) can be conveniently applied as parallel lines.

[0085] The LiPA technique, as described by Stuyver et al. (1993) and in international application WO 94/12670, provides a very rapid and user-friendly hybridization test. Results can be read within 4 h. after the start of the amplification. After amplification during which usually a non-isotopic label is incorporated in the amplified product, and alkaline denaturation, the amplified product is contacted with the probes on the membrane and the hybridization is carried out for about 1 to 1.5 h. Consequently, the hybrids formed are detected by an enzymatic procedure resulting in a visual purple-brown precipitate. The LiPA format is completely compatible with commercially available scanning devices, thus rendering automatic interpretation of the results possible. All those advantages make the LiPA format liable for use in a routine setting.

[0086] The LiPA format is not only an advantageous tool for identification and detection of pathogens at the species level but also at higher or lower taxonomical levels. For instance, probe-configurations on LiPA strips can be selected in such a manner that they can detect a complete genus (e.g. Neisseria, Listeria, etc.) or can identify subgroups within a genus (e.g. subgroups in the Mycobacterium avium-intracellulare-scorfulaceum complex) or can in some cases even detect subtypes (subspecies, serovars, sequevars, biovars, etc. whatever is clinically relevant) within a species.

[0087] It should be stressed that the ability to simultaneously generate hybridization results with a number of probes is an outstanding benefit of the LiPA technology. In many cases the amount of information which can be obtained by a particular combination of probes greatly outnumbers the data obtained by using single probe assays. Therefor the selection of probes on the membrane strip is of utmost importance since an optimized set of probes will generate the maximum of information possible. This is more particularly exemplified further herein.

[0088] The fact that different probes can be combined on one strip also offers the possibility to conveniently cope

with a lack of sensitivity due to sequence heterogeneity in the target region of the group of organisms to be detected. Due to this heterogeneity, two or more probes may be required to positively identify all organisms of the particular group. These probes can be applied to membrane strips at different locations and the result is interpreted as positive if at least one of these probes is positive. Alternatively these probes can be applied as a mixture at the same location, thereby reducing the number of lines on a strip. This reduction may be convenient in order to make the strip more concise or to be able to extend the total number of probes on one strip. Another alternative approach, in view of its practical benefits, is the synthesis of oligonucleotides harbouring the sequences of two (or more) different probes (=degenerate probes) which then can be further processed and applied to the strip as one oligonucleotide molecule. This approach would considerably simplify the manufacturing procedures of the LiPA-strips. For example, probes with nucleotide sequences A and B are both required to detect all strains of taxon X. In the latter alternative a probe can be synthesized having the nucleotide sequence AB. This probe will have the combined characteristics of probes A and B.

[0089] By virtue of the above-mentioned properties the LiPA system can be considered as a preferential format for a hybridization method wherein several organisms need to be detected simultaneously in a sample. Moreover, as described in the examples section, the LiPA system is a preferred format for a selection method for the experimental evaluation and selection of theoretically designed probes.

[0090] However, it should be clear that any other hybridization assay, whereby different probes are used under the same hybridization and wash conditions can be used for the above-mentioned detection and/or selection methods. For example, it may be possible to immobilize the target nucleic acid to a solid support, and use mixtures of different probes, all differently labeled, resulting in a different detection signal for each of the probes hybridized to the target.

[0091] As an example, the procedure to be followed for the detection of one or more organisms in a sample using the LiPA format is outlined below :

- First, the nucleic acids of the organism(s) to be detected in the sample, is made available for amplification and/or hybridization.
- Secondly, the nucleic acids, if present, are amplified with one or another target amplification system, as specified below. Usually, amplification is needed to enhance the subsequent hybridization signal. However for some samples or some organisms amplification might not be necessary. This might also be the case if, for the detection of the hybrids formed, highly sensitive signal-amplification systems are used.
- Thirdly, eventually after a denaturation step, the nucleic acids present in the sample or the resulting amplified product are contacted with LiPA strips onto which one or more DNA-probes, allowing the detection of the organisms of interest, are immobilized, and hybridization is allowed to proceed.
- Finally, eventually after having performed a wash step, the hybrids are detected using a convenient and compatible detection system. From the hybridization signals or patterns observed the presence or absence of one or several organisms screened for in that particular biological sample can be deduced.

[0092] The amplification system used may be more or less universal, depending on the specific application needed.

[0093] By using universal primers located in the conserved flanking regions (16S and 23S gene) of the rRNA spacer, the spacer region of most if not all organisms of eubacterial origin will be amplified. The same result may be obtained by using a combination of different sets of primers with reduced universality (multiplex amplification, i.e. an amplification procedure in which two or more primer sets are used simultaneously in one and the same reaction mixture).

[0094] For some applications it may be appropriate to amplify not all organisms present in the sample but more specifically, beforehand defined taxa. This may be achieved using specific primers located either in less conserved parts of the flanking genes of the spacers (e.g. MYCP1-5 for amplification of the spacer region of mycobacteria) or located in the spacers themselves (e.g. LIS-P1-P7 , BRU-P1-4, CHTR-P1-2 and YEC-P1-2 for specific amplification of the spacer region(s) of Listeria species, Brucella species, Chlamydia trachomatis, and Yersinia enterocolitica respectively).

[0095] The present invention thus provides a method for detection and identification of at least one micro-organism, or for the simultaneous detection of several micro-organisms in a sample, comprising the steps of:

- 50 (i) if need be releasing, isolating and/or concentrating the polynucleic acids from the micro-organism(s) to be detected in the sample;
- (ii) if need be amplifying the 16S-23S rRNA spacer region, or a part of it, from the micro-organism(s) to be detected, with at least one suitable primer pair;
- 55 (iii) hybridizing the polynucleic acids of step (i) or (ii) with a set of probes comprising at least two probes, under the same hybridization and wash conditions, with said probes being selected from the sequences of table 1a or equivalents thereof and/or from taxon-specific probes derived from any of the spacer sequences represented in figs. 1-103, with said taxon-specific probe being selected such that it is capable of hybridizing under the same

hybridization and wash conditions as at least one of the probes of table 1a;
 (iv) detecting the hybrids formed in step (iii);
 (v) identification of the micro-organism(s) present in the sample from the differential hybridization signals obtained in step (iv).

5 [0096] The probes as mentioned in table 1a are all selected in such a way that they show the desired hybridization characteristics at a hybridization and wash temperature of 50°C in a preferred hybridization and wash medium of 3X SSC and 20% formamide.

10 [0097] The term "equivalents" of a probe, also called "variants" or "homologues" or "obvious derivatives", refers to probes differing in sequence from any of the probes specified in table 1 either by addition to or removal from any of their respective extremities of one or several nucleotides, or by changing one or more nucleotides within said sequences, or a combination of both, provided that said equivalents still hybridize with the same RNA or DNA target as the corresponding unmodified probe sequence. Said equivalents share at least 75% homology, preferably more than 80%, most preferably more than 85% homology with the corresponding unmodified probe sequence. It should be noted that, 15 when using an equivalent of a probe, it may be necessary to modify the hybridization conditions to obtain the same specificity as the corresponding unmodified probe. As a consequence, since it is the aim of this invention to use a set of probes which work under the same hybridization and wash conditions, it will also be necessary to modify accordingly the sequence of the other probes, belonging to the same set as the original unmodified probe. These modifications can be done according to principles known in the art, e.g. such as those described in Hames B and Higgins S (Eds): 20 Nucleic acid hybridization. Practical approach. IRL Press, Oxford, UK, 1985.

[0098] The invention also provides for a method to select taxon-specific probes from the spacer region sequence(s) of said taxon, said probes being selected such that they show their desired hybridization characteristics under unified hybridization and wash conditions.

[0099] The term "unified" conditions means that these conditions are the same for the different probes enabling the detection of different taxa.

[0100] Preferentially, the present invention provides for a method as described above wherein at least 2 micro-organisms are detected simultaneously.

[0101] In a preferred embodiment, the set of probes as described in step (iii) is comprising at least two probes being selected from the sequences of table 1a, or equivalents thereof.

30 [0102] In another embodiment, the set of probes as described in step (iii) is comprising at least one probe being selected from the sequences of table 1a, or equivalents thereof, and at least one taxon-specific probe derived from any of the spacer sequences as represented in figs. 1-103.

[0103] In still another embodiment, the set of probes as described in step (iii) is comprising at least two taxon-specific probes derived from any of the spacer sequences as represented in figs. 1-103.

35 [0104] The present invention also provides for a method as described above, wherein the probes as specified in step (iii) are combined with at least one other probe, preferentially also from the 16S-23S rRNA spacer region, enabling the simultaneous detection of different pathogenic bacteria liable to be present in the same sample.

[0105] The organisms of clinical relevance present in biological samples may vary considerably depending on the origin of the sample. The most common pathogenic bacteria which may be found in sputum samples, or in samples 40 originating from the respiratory tract, are :

- Moraxella catarrhalis
- Streptococcus pneumoniae
- Haemophilus influenzae
- Pseudomonas aeruginosa
- Mycoplasma pneumoniae
- Acinetobacter species
- Mycobacterium species
- Staphylococcus aureus
- Legionella pneumophila

[0106] A LiPA-strip harbouring spacer-probes enabling the detection of most if not all of these organisms would be extremely beneficial for reasons explained above.

[0107] Evidently, this also applies for other biological samples, as there are :

55 cerebrospinal fluid, urogenital samples, gastrointestinal samples, blood, urine, food products, soil, etc. For example, a preferred panel for cerebrospinal fluid would contain probe combinations enabling the detection and differentiation of the following organisms :

- Neisseria meningitidis
 - Streptococcus pneumoniae
 - Streptococcus agalactiae
 - Listeria monocytogenes
 - Mycobacterium tuberculosis

[0108] For some of the above mentioned organisms, spacer probes were already designed in a previous patent application (WO 91/16454). In order to be able to detect most pathogens possibly present in a sample in a single test, the probes of the present invention may be combined with at least one of the probes of WO 91/16454, or their obvious derivatives as specified in WO 91/16454. For clarity, these probes are listed hereafter:

Neisseria gonorrhoeae: NGI1: CGATGCGTCGTTATTCTACTTCCG

15 NGI2: TTCGTTTACCTACCCGTTGACTAAGTAAGCAAAAC

Neisseria meningitidis: NM11: GGTCAAGTGTGACGTGGCCCTG

20 NMII2: GTTCTTGGTCAAGTGTGACGTC

NMI3: GCGTTCGTTATAGCTATCTACTGTGC

NM14: TGC GTTCGATATTGCTATCTACTGTGCA

NM15: TTTTGTCTTGGTCAAGTGTGACGTCCC

25 NM15: TTTGTTCTGGTCAAGTGTGACGTCGCCCTGAA
TGCGATTGCTGGATTC

TGGATTCTGTTCCATT

NMI6: TTTGCCTAACATTCCGTTGACTAGAACATCAGAC

³⁰ Haemophilus ducreyi HDI1: TTATTATGCGCGAGGCATATTG

Branhamella catharralis BCI1: TTAAACATCTTACCAAAG

BCI2: TTGATGTTAAACTTGCTT

BP11: CCACACCCATCCTCTGGACACGCCCTT

Haemophilus influenzae HJ11: ACGCATGAAATTGACCCCCACTT

III2: -ACTTCTGAACTGAAAAGTTAAAAC

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SAIZ. JUAAACCIACCATTGCTT

SAIS: TCCACCGATCTAGAAAATAGATCTGTAGAA

SAI4: TCTAGTTTAAAGAACTAGG

Streptococcus pneumoniae SPII: GTGAGAGATCACCAAGTAATGCA

50 SPI2: AGGAAC TGCGCAT TGGTCTT

SPI3: GAGTTATGACTGAAAGGTCAAA

[0109] The invention thus provides for a method as described above, wherein said sample is originating from the respiratory tract, and wherein the set of probes as defined in step (iii) comprises at least one probe chosen from the following spacer probes:

MYC-ICG-1 :	ACTGGATAGTGGTTGCGAGCATCTA	(SEQ ID NO 1)
MYC-ICG-22 :	CTTCTGAATAGTGGTTGCGAGCATCT	(SEQ ID NO 2)
5 MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGGCCA	(SEQ ID NO 3)
MTB-ICG-2 :	GACTTGTCCAGGTGTTGCCAC	(SEQ ID NO 4)
MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)
10 MAI-ICG-1 :	CAACAGCAAATGATTGCCAGACACAC	(SEQ ID NO 6)
MIL-ICG-11 :	GAGGGGTTCCCGTCTGTAGTG	(SEQ ID NO 7)
15 MIL-ICG-22 :	TGAGGGGTTCTCGTCTGTAGTG	(SEQ ID NO 8)
MAC-ICG-1 :	CACTCGGTCGATCCGTGTGGA	(SEQ ID NO 9)
MAV-ICG-1 :	TCGGTCCGTCCGTGTGGAGTC	(SEQ ID NO 10)

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	MAV-ICG-22 : GTGGCCGGCGTCATCGAAA	(SEQ ID NO 11)
5	MIN-ICG-1 : GCATAGTCCTAGGGCTGATGCGTT	(SEQ ID NO 12)
	MIN-ICG-2 : GCTGATGCGTCGTCGAAATGTGT	(SEQ ID NO 13)
	MIN-ICG-22 : CTGATGCGTCGTCGAAATGTGT	(SEQ ID NO 14)
	MIN-ICG-222 : TGATGCGTCGTCGAAATGTGT	(SEQ ID NO 15)
10	MIN-ICG-2222 : GGCTGATGCGTCGTCGAAATGTGTAA	(SEQ ID NO 16)
	MAL-ICG-1 : ACTAGATGAACGCGTAGTCCTTGT	(SEQ ID NO 17)
	MHEF-ICG-1 : TGGACGAAAACCGGGTGCACAA	(SEQ ID NO 18)
15	MAH-ICG-1 : GTGTAATTCTTTAACTCTGTGTAAAGTAAGTG	(SEQ ID NO 19)
	MCO-ICG-11 : TGGCCGGCGTGTTCATCGAAA	(SEQ ID NO 20)
20	MTH-ICG-11 : GCACTTCAATTGGTGAAGTGCGAGCC	(SEQ ID NO 21)
	MTH-ICG-2 : GCGTGGTCTTCATGCCCG	(SEQ ID NO 22)
	MEF-ICG-11 : ACGCGTGGTCCTCGTGG	(SEQ ID NO 23)
25	MSC-ICG-1 : TCGGCTCGTTCTGAGTGGTGT	(SEQ ID NO 24)
	MKA-ICG-1 : GATGCGTTGCTACGGGTAGCGT	(SEQ ID NO 25)
	MKA-ICG-2 : GATGCGTTGCCTACGGGTAGCGT	(SEQ ID NO 26)
30	MKA-ICG-3 : ATGCGTTGCCCTACGGGTAGCGT	(SEQ ID NO 27)
	MKA-ICG-4 : CGGGCTCTGTCGAGAGTTGTC	(SEQ ID NO 28)
	MKA-ICG-5 : CCCTCAGGGATTTCTGGGTGTG	(SEQ ID NO 182)
35	MKA-ICG-6 : GGACTCGTCCAAGAGTGTGTCC	(SEQ ID NO 183)
	MKA-ICG-7 : TCGGGCTTGGCCAGAGCTGT	(SEQ ID NO 184)
	MKA-ICG-8 : GGGTGCACAGCAAGCGA	(SEQ ID NO 185)
40	MKA-ICG-9 : GATGCGTTGCCCTACGGG	(SEQ ID NO 186)
	MKA-ICG-10 : CCCTACGGGTAGCGTGTCTTTG	(SEQ ID NO 187)
	MCH-ICG-1 : GGTGTGGACTTGACTTCTGAATAG	(SEQ ID NO 29)
45	MCH-ICG-2 : CGGCAAAACGTCGGACTGTCA	(SEQ ID NO 30)
	MCH-ICG-3 : GGTGTGGTCCTGACTTATGGATAG	(SEQ ID NO 210)
	MGO-ICG-1 : AACACCCCTCGGGTGCTGTCC	(SEQ ID NO 31)
50	MGO-ICG-2 : GTATGCGTTGTCGTCGCC	(SEQ ID NO 32)
	MGO-ICG-5 : CGTGAGGGTCATCGTCTGTAG	(SEQ ID NO 33)
	MUL-ICG-1 : GGTTTCGGGATGTTGTCCCACC	(SEQ ID NO 175)
55	MGV-ICG-1 : CGACTGAGGTGACGTGGTGT	(SEQ ID NO 176)

	MGV-ICG-2 :	GGTGTGAGCATTGAATAGTGGTTGC	(SEQ ID NO 177)
	MGV-ICG-3 :	TCGGGCCGCGTGGTCGTCAAA	(SEQ ID NO 211)
5	MXE-ICG-1 :	GTTGGCAGCAGGCAGTAACC	(SEQ ID NO 178)
	MSI-ICG-1 :	CCGGCAACGGTTACGTGTTC	(SEQ ID NO 179)
	MFO-ICG-1 :	TCGTTGGATGGCCTCGCACCT	(SEQ ID NO 180)
10	MFO-ICG-2 :	ACTTGGCGTGGGATGCGGGAA	(SEQ ID NO 181)
	MML-ICG-1 :	CGGATCGATTGAGTGCTTGTCCC	(SEQ ID NO 188)
15	MML-ICG-2 :	TCTAAATGAACGCAGTGCCTGG	(SEQ ID NO 189)
	MCE-ICG-1 :	TGAGGGAGCCCGTGCCTGTA	(SEQ ID NO 190)
	MHP-ICG-1 :	CATGTTGGGCTTGATCGGGTGC	(SEQ ID NO 191)
20	PA-ICG 1 :	TGGTGTGCTGCGTGATCCGAT	(SEQ ID NO 34)
	PA-ICG 2 :	TGAATGTTCTGGATGAAACATTGATT	(SEQ ID NO 35)
	PA-ICG 3 :	CACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 36)
25	PA-ICG 4 :	TGAATGTTCTG(G/A)(G/A)ATGAACATTGATTCTGGTC	(SEQ ID NO 37)
	PA-ICG 5 :	CTCTTCACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 38)
30	MPN-ICG 1 :	ATCGGTGGTAAATTAAACCCAAATCCCTGT	(SEQ ID NO 49)
	MPN-ICG 2 :	CAGTTCTGAAAGAACATTCCGCTTCTTC	(SEQ ID NO 50)
	MGE-ICG 1 :	CACCCATTAATTTTCGGTGTAAAACCC	(SEQ ID NO 51)
35	Mycoplasma-ICG :	CAAAACTGAAAACGACAATCTTCTAGTTCC	(SEQ ID NO 52)
	STAU-ICG 1 :	TACCAAGCAAAACCGAGTGAATAAAGAGTT	(SEQ ID NO 53)
	STAU-ICG 2 :	CAGAAGATGCGGAATAACGTGAC	(SEQ ID NO 54)
40	STAU-ICG 3 :	AACGAAGCCGTATGTGAGCATTGAC	(SEQ ID NO 55)
	STAU-ICG 4 :	GAACGTAACCTCATGTTAACGTTGACTTAT	(SEQ ID NO 56)
	ACI-ICG 1 :	GCTTAAGTGCACAGTGCTCTAAACTGA	(SEQ ID NO 57)
45	ACI-ICG 2 :	CACGGTAATTAGTGTGATCTGACGAAG	(SEQ ID NO 58)

and more preferably from the following spacer probes:

MYC-ICG-1 :	ACTGGATAGTGGTTGCGAGCATCTA	(SEQ ID NO 1)
5 MYC-ICG-22 :	CTTCTGAATAGTGGTTGCGAGCATCT	(SEQ ID NO 2)
MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGGCCA	(SEQ ID NO 3)
10 MTB-ICG-2 :	GACTTGTCCAGGTGTTGTCCCAC	(SEQ ID NO 4)
MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)
15 MAI-ICG-1 :	CAACAGCAAATGATTGCCAGACACAC	(SEQ ID NO 6)

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	MIL-ICG-11 :	GAGGGGTTCCCGTCTGTAGTG	(SEQ ID NO 7)
	MIL-ICG-22 :	TGAGGGGTTCTCGTCTGTAGTG	(SEQ ID NO 8)
5	MAC-ICG-1 :	CACTCGGTGATCCGTGTGGA	(SEQ ID NO 9)
	MAV-ICG-1 :	TCGGTCCCGTCCGTGTGGAGTC	(SEQ ID NO 10)
	MAV-ICG-22 :	GTGCCCGCGTTCATCGAAA	(SEQ ID NO 11)
10	MIN-ICG-1 :	GCATAGTCCTTAGGGCTGATGCGTT	(SEQ ID NO 12)
	MAL-ICG-1 :	ACTAGATGAACCGTAGCCTTGT	(SEQ ID NO 17)
	MCO-ICG-11 :	TGGCCGGCGTGTTCATCGAAA	(SEQ ID NO 20)
15	MTH-ICG-11 :	GCACTTCAATTGGTGAAGTGCAGGCC	(SEQ ID NO 21)
	MTH-ICG-2 :	GCGTGGTCTTCATGCCGG	(SEQ ID NO 22)
	MEF-ICG-11 :	ACGCGTGGTCCTCGTGG	(SEQ ID NO 23)
20	MSC-ICG-1 :	TCGGCTCGTCTGAGTGGTGTGTC	(SEQ ID NO 24)
	MKA-ICG-3 :	ATGCGTTGCCCTACGGTAGCGT	(SEQ ID NO 27)
	MKA-ICG-4 :	CGGGCTCTGTTGAGAGAGTTGTC	(SEQ ID NO 28)
25	MKA-ICG-5 :	CCCTCAGGGATTTCTGGGTGTG	(SEQ ID NO 182)
	MKA-ICG-6 :	GGACTCGTCCAAGAGTGTGTGTC	(SEQ ID NO 183)
	MKA-ICG-7 :	TCGGGCTTGGCCAGAGCTGTT	(SEQ ID NO 184)
30	MKA-ICG-8 :	GGGTGCGAACAGCAAGCGA	(SEQ ID NO 185)
	MKA-ICG-9 :	GATGCGTTGCCCTACGGG	(SEQ ID NO 186)
	MKA-ICG-10 :	CCCTACGGTAGCGTGTCTTTG	(SEQ ID NO 187)
35	MCH-ICG-1 :	GGTGTGGACTTGACTTCTGAATAG	(SEQ ID NO 29)
	MCH-ICG-2 :	CGGCAAAACGTGGACTGTCA	(SEQ ID NO 30)
	MCH-ICG-3 :	GGTGTGGCCTTGACTTATGGATAG	(SEQ ID NO 210)
40	MGO-ICG-5 :	CGTGAGGGTCATCGTCTGTAG	(SEQ ID NO 33)
	MUL-ICG-1 :	GGTTTCGGGATGTTGTCCCACC	(SEQ ID NO 175)
	MGV-ICG-1 :	CGACTGAGGTGACGTGGTGT	(SEQ ID NO 176)
45	MGV-ICG-2 :	GGTGTGGAGCATTGAATAGTGGTTGC	(SEQ ID NO 177)
	MGV-ICG-3 :	TCGGGCCGCGTGTTCGTCAA	(SEQ ID NO 211)
	MXE-ICG-1 :	GTTGGGCAGCAGGCAGTAACC	(SEQ ID NO 178)
50	MSI-ICG-1 :	CCGGCAACGGTTACGTGTC	(SEQ ID NO 179)
	MFO-ICG-1 :	TCGTTGGATGCCCTCGCACCT	(SEQ ID NO 180)
	MFO-ICG-2 :	ACTTGGCGTGGGATGCGGGAA	(SEQ ID NO 181)
55	MML-ICG-1 :	CGGATCGATTGAGTGCTTGTCCC	(SEQ ID NO 188)

MML-ICG-2 :	TCTAAATGAACGCAGTGGCGATGG	(SEQ ID NO 189)
MCE-ICG-1 :	TGAGGGAGCCCGTGCGCTGTA	(SEQ ID NO 190)
5 MHP-ICG-1 :	CATGTTGGGCTGATCGGGTGC	(SEQ ID NO 191)
PA-ICG 1 :	TGGTGTGCTGCGTGATCCGAT	(SEQ ID NO 34)
10 PA-ICG 4 :	TGAATGTTCGT(G/A)(G/A)ATGAACATTGATTCTGGTC	(SEQ ID NO 37)
15 PA-ICG 5 :	CTCTTCACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 38)
MPN-ICG 1 :	ATCGGTGGTAAATTAAACCAAATCCCTGT	(SEQ ID NO 49)
MPN-ICG 2 :	CAGTTCTGAAAGAACATTCCGCTCTTC	(SEQ ID NO 50)
20 MGE-ICG 1 :	CACCCATTAATTTTCGGTGTAAAACCC	(SEQ ID NO 51)
Mycoplasma-ICG :	CAAAACTGAAAACGACAATCTTCTAGTTCC	(SEQ ID NO 52)
STAU-ICG 1 :	TACCAAGCAAAACCGAGTGAATAAAGAGTT	(SEQ ID NO 53)
25 STAU-ICG 2 :	CAGAAGATGCCGAATAACGTGAC	(SEQ ID NO 54)
STAU-ICG 3 :	AACGAAGCCGTATGTGAGCATTGAC	(SEQ ID NO 55)
STAU-ICG 4 :	GAACGTAACCATGTTAACGTTGACTTAT	(SEQ ID NO 56)
30 ACI-ICG 1 :	GCTTAAGTGCACAGTGCTCTAAACTGA	(SEQ ID NO 57)
ACI-ICG 2 :	CACGGTAATTAGTGTGATCTGACGAAG	(SEQ ID NO 58)

or equivalents of said probes,

35 and/or wherein the set of probes comprises at least one taxon-specific probe derived from the spacer region sequence corresponding to one of the micro-organisms to be detected in said sample, said spacer region sequence being chosen from any of the sequences as represented by SEQ ID NO 76 to 106, 157 to 174, 124, 125, 111 to 115, 139 to 144, or 126 to 130,
40 and with said probes or equivalents being possibly used in combination with any probe detecting at least one of the following organisms: Haemophilus influenzae, Streptococcus pneumoniae, Moraxella catarrhalis or Bordetella pertussis.

[0110] The above mentioned probes of the invention are designed for the detection of Mycobacterium species (SEQ ID NO 1 to 33 and 175 to 191), of Pseudomonas aeruginosa (SEQ ID NO 34 to 38), of Mycoplasma species (SEQ ID NO 49 to 52), of Staphylococcus aureus (SEQ ID NO 53 to 56) and of Acinetobacter baumanii (SEQ ID NO 57 and 58).

45 [0111] Preferentially, at least two, three, four, five, six, seven, eight or more of said probes are used simultaneously.
[0112] The invention also relates to a method as described above, wherein said sample is a sample taken from the cerebrospinal fluid, and wherein the set of probes as described in step (iii) comprises at least one probe chosen from the following spacer probes:

MYC-ICG-1 :	ACTGGATAGTGGTTGCGAGCATCTA	(SEQ ID NO 1)
MYC-ICG-22 :	CTTCTGAATAGTGGTTGCGAGCATCT	(SEQ ID NO 2)
5 MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGGCCA	(SEQ ID NO 3)
MTB-ICG-2 :	GACTTGTCCAGGTGTTGTCCCAC	(SEQ ID NO 4)
10 MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)
LIS-ICG 1 :	CAAGTAACCGAGAATCATCTGAAAGTGAATC	(SEQ ID NO 39)
LMO-ICG 1 :	AAACAAACCTTACTTCGTAGAAGTAAATTGGTTAAG	
15		(SEQ ID NO 40)
LMO-ICG 2 :	TGAGAGGTTAGTACTTCTCAGTATGTTGTT	(SEQ ID NO 41)
LMO-ICG 3 :	AGGCACTATGCTGAAGCATCGC	(SEQ ID NO 42)
20 LISP-ICG 1:	CGTTTCATAAGCGATCGCACGTT	(SEQ ID NO 212)

and preferably from the following spacer probes:

25 MYC-ICG-1 :	ACTGGATAGTGGTTGCGAGCATCTA	(SEQ ID NO 1)
MYC-ICG-22 :	CTTCTGAATAGTGGTTGCGAGCATCT	(SEQ ID NO 2)
MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGGCCA	(SEQ ID NO 3)
30 MTB-ICG-2 :	GACTTGTCCAGGTGTTGTCCCAC	(SEQ ID NO 4)
MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)
35 LIS-ICG 1 :	CAAGTAACCGAGAATCATCTGAAAGTGAATC	(SEQ ID NO 39)
LMO-ICG 3 :	AGGCACTATGCTGAAGCATCGC	(SEQ ID NO 42)
40 LISP-ICG 1:	CGTTTCATAAGCGATCGCACGTT	(SEQ ID NO 212)

or equivalents of said probes,

and/or wherein the set of probes comprises at least one taxon-specific probe derived from the spacer region sequence corresponding to one of the micro-organisms to be detected in said sample, said spacer region sequence being chosen

45 from any of the sequences as represented by SEQ ID NO 116, 118-121, or 213-215,
and with said probes or equivalents being possibly used in combination with any probe detecting at least one of the following organisms: Neisseria meningitidis, Haemophilus influenzae or Streptococcus pneumoniae.

50 [0113] The above mentioned probes of the invention are designed for the detection of Mycobacterium species, and more particularly Mycobacterium tuberculosis (SEQ ID NO 1 to 5), and of Listeria species, more particularly Listeria monocytogenes (SEQ ID NO 39 to 42).

[0114] Preferentially, at least two, three, four, five, six, seven, eight or more of said probes are used simultaneously.

[0115] The invention also relates to a method as described above, wherein said sample is a sample taken from the urogenital tract, and wherein the set of probes as described in step (iii) comprises at least one probe chosen from the following spacer probes:

CHTR-ICG 1 :	GGAAGAACGCTGAGAAGGTTCTGAC	(SEQ ID NO 45)
CHTR-ICG 2 :	GCATTATATGTAAGAGCAAGCATTCTATTCA	(SEQ ID NO 46)
5 CHTR-ICG 3 :	GAGTAGCGTGGTGAGGACGAGA	(SEQ ID NO 47)
CHTR-ICG 4 :	GAGTAGCGCGGTGAGGACGAGA	(SEQ ID NO 201)
10 CHPS-ICG 1 :	GGATAACTGTCTTAGGACGGTTGAC	(SEQ ID NO 48)
MGE-ICG 1 :	CACCCATTAATTTTCGGTGTAAAACCC	(SEQ ID NO 51)
Mycoplasma-ICG :	CAAAACTGAAAACGACAATCTTCTAGTTCC	(SEQ ID NO 52)

15 or equivalents of said probes,
and/or wherein the set of probes comprises at least one taxon-specific probe derived from the spacer region sequence
corresponding to one of the micro-organisms to be detected in said sample, said spacer region sequence being chosen
from any of the sequences as represented by SEQ ID NO 122, 123, 197, 124 or 125,
with said probes or equivalents being possibly used in combination with any probe detecting at least one of the following
20 organisms: Neisseria gonorrhoeae, Haemophilus ducreyi or Streptococcus agalactiae.
[0116] The above mentioned probes of the invention are designed for the detection of Chlamydia species (SEQ ID
NO 45 to 48 and 201) and of Mycoplasma species (SEQ ID NO 51 and 52).
[0117] Preferentially, at least two, three, four, five, six or seven of said probes are used simultaneously.
[0118] The invention also relates to a method as described above, wherein said sample is a sample taken from food,
25 and wherein the set of probes as defined in step (iii) comprises at least one probe chosen from the following spacer
probes:

LIS-ICG 1 :	CAAGTAACCGAGAATCATCTGAAAGTGAATC	(SEQ ID NO 39)
30 LMO-ICG 1 :	AAACAAACCTTACTTCGTAGAAGTAAATTGGTTAAG	
		(SEQ ID NO 40)
LMO-ICG 2 :	TGAGAGGTTAGTACTTCTCAGTATGTTGTT	(SEQ ID NO 41)

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LMO-ICG 3 :	AGGCACATATGCTTGAAGCATCGC	(SEQ ID NO 42)	
LIV-ICG 1 :	GTTAGCATAAAATAGGTAACTATTATGACACAAGTAAC		
5		(SEQ ID NO 43)	
LSE-ICG 1 :	AGTTAGCATAAGTAGTGTAACATTATGACACAAG	(SEQ ID NO 44)	
10	LISP-ICG 1:	CGTTTCATAAGCGATCGCACGTT	(SEQ ID NO 212)
STAU-ICG 1 :	TACCAAGCAAAACCGAGTGAATAAAGAGTT	(SEQ ID NO 53)	
15	STAU-ICG 2 :	CAGAACGATGCGGAATAACGTGAC	(SEQ ID NO 54)
STAU-ICG 3 :	AACGAAGCCGTATGTGAGCATTGAC	(SEQ ID NO 55)	
20	STAU-ICG 4 :	GAACGTAACCTCATGTTAACGTTGACTTAT	(SEQ ID NO 56)
BRU-ICG 1 :	CGTGCCGCCTCGTTCTCTTT	(SEQ ID NO 59)	
25	BRU-ICG 2 :	TTCGCTTCGGGGTGGATCTGTG	(SEQ ID NO 60)
BRU-ICG 3 :	GCGTAGTAGCGTTGCGTCGG	(SEQ ID NO 193)	
BRU-ICG 4 :	CGCAAGAACGCTTGCTCAAGCC	(SEQ ID NO 194)	
30	SALM-ICG 1 :	CAAAACTGACTTACGAGTCACGTTGAG	(SEQ ID NO 61)
SALM-ICG 2 :	GATGTATGCTCGTTATTCCACGCC	(SEQ ID NO 62)	
STY-ICG 1 :	GGTCAAACCTCCAGGGACGCC	(SEQ ID NO 63)	
35	SED-ICG 1 :	GCGGTAATGTGTGAAAGCGTTGCC	(SEQ ID NO 64)
YEC-ICG 1 :	GGAAAAGGTACTGCACGTGACTG	(SEQ ID NO 198)	
YEC-ICG 2 :	GACAGCTGAAACTTATCCCTCCG	(SEQ ID NO 199)	
YEC-ICG 3 :	GCTACCTGTTGATGTAATGAGTCAC	(SEQ ID NO 200)	

and preferably from the following spacer probes:

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LIS-ICG 1 :	CAAGTAACCGAGAATCATCTGAAAGTGAATC	(SEQ ID NO 39)
LMO-ICG 3 :	AGGCACTATGCTTGAAGCATCGC	(SEQ ID NO 42)
5 LISP-ICG 1:	CGTTTCATAAGCGATCGCACGTT	(SEQ ID NO 212)
STAU-ICG 1 :	TACCAAGCAAAACCGAGTGAATAAAGAGTT	(SEQ ID NO 53)
10 STAU-ICG 2 :	CAGAAGATGCGGAATAACGTGAC	(SEQ ID NO 54)
STAU-ICG 3 :	AACGAAGCCGTATGTGAGCATTGAC	(SEQ ID NO 55)
15 STAU-ICG 4 :	GAACGTAACCTCATGTTAACGTTGACTTAT	(SEQ ID NO 56)
BRU-ICG 2 :	TTCGCTTCGGGGTGGATCTGTG	(SEQ ID NO 60)
BRU-ICG 3 :	GCGTAGTAGCGTTGCGTCGG	(SEQ ID NO 193)
BRU-ICG 4 :	CGCAAGAAGCTTGCTCAAGCC	(SEQ ID NO 194)
20 SALM-ICG 1 :	CAAAACTGACTTACGAGTCACGTTGAG	(SEQ ID NO 61)
YEC-ICG 1 :	GGAAAAGGTACTGCACGTGACTG	(SEQ ID NO 198)
25 YEC-ICG 2 :	GACAGCTGAAACTTATCCCTCCG	(SEQ ID NO 199)
YEC-ICG 3 :	GCTACCTGTTGATGTAATGAGTCAC	(SEQ ID NO 200)

or equivalents of said probes,

30 and/or wherein the set of probes comprises at least one taxon-specific probe derived from the spacer region sequence corresponding to one of the micro-organisms to be detected in said sample, said spacer region sequence being chosen from any of the sequences as represented by SEQ ID NO 116, 118 -121,213-215, 139-144, 131, 132, 154, 133-138, 195 or 196,

35 with said probes or equivalents being possibly used in combination with any probe detecting strains of Campylobacter species.

[0119] The above mentioned probes of the invention are designed for the detection of Listeria species (SEQ ID NO 39 to 44), of Staphylococcus species (SEQ ID NO 53 to 56), of Brucella species (SEQ ID NO 59, 60, 193 and 194), of Salmonella species (SEQ ID NO 61 to 64) and of Yersinia enterocolitica (SEQ ID NO 198 to 200).

[0120] Preferentially, at least two, three, four, five, six, seven, eight or more of said probes are used simultaneously.

40 [0121] The invention also relates to a method as described above, wherein said sample is a sample taken from the gastrointestinal tract of a patient, and wherein the set of probes as defined in step (iii) comprises at least one probe chosen from the following spacer probes:

45 SALM-ICG 1 :	CAAAACTGACTTACGAGTCACGTTGAG	(SEQ ID NO 61)
SALM-ICG 2 :	GATGTATGCTTCGTTATTCCACGCC	(SEQ ID NO 62)
STY-ICG 1 :	GGTCAAACCTCCAGGGACGCC	(SEQ ID NO 63)
50 SED-ICG 1 :	GCGGTAATGTGTGAAAGCGTTGCC	(SEQ ID NO 64)
YEC-ICG 1 :	GGAAAAGGTACTGCACGTGACTG	(SEQ ID NO 198)
YEC-ICG 2 :	GACAGCTGAAACTTATCCCTCCG	(SEQ ID NO 199)
55 YEC-ICG 3 :	GCTACCTGTTGATGTAATGAGTCAC	(SEQ ID NO 200)

and preferably from the following spacer probes:

SALM-ICG 1 :	CAAAACTGACTTACGAGTCACGTTGAG	(SEQ ID NO 61)
YEC-ICG 1 :	GGAAAAGGTACTGCACGTGACTG	(SEQ ID NO 198)
5 YEC-ICG 2 :	GACAGCTGAAACTTATCCCTCCG	(SEQ ID NO 199)
YEC-ICG 3 :	GCTACCTGTTGATGTAATGAGTCAC	(SEQ ID NO 200)

- 10 or equivalents of said probes,
and/or wherein the set of probes comprises at least one taxon-specific probe derived from the spacer region sequence
corresponding to one of the micro-organisms to be detected in said sample, said spacer region sequence being chosen
from any of the sequences as represented by SEQ ID NO 133-138 or 195-196,
with said probes or equivalents being possibly used in combination with any probe detecting Campylobacter species.
15 [0122] The above mentioned probes of the invention are designed to detect Salmonella species (SEQ ID NO 61 to
64) and Yersinia enterocolitica (SEQ ID NO 198 to 200).
[0123] Preferentially, at least two, three, four, five, six or seven of said probes are used simultaneously.
[0124] The invention also relates to the use of the selected probes or their equivalents for the detection of specific
20 bacterial taxa, said taxa being either a complete genus, or a subgroup within a genus, a species, or even a subtype
within a species.
[0125] The invention thus provides for a method as described above to detect and identify one or more strains of
Mycobacterium species and subspecies in a sample, wherein step (iii) comprises hybridizing to at least one of the
following probes:

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MYC-ICG-1 :	ACTGGATAGTGGTTGCGAGCATCTA	(SEQ ID NO 1)
MYC-ICG-22 :	CTTCTGAATAGTGGTTGCGAGCATCT	(SEQ ID NO 2)
5 MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGGCCA	(SEQ ID NO 3)
MTB-ICG-2 :	GACTTGTCCAGGTGTTGTCCCAC	(SEQ ID NO 4)
10 MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)
MAI-ICG-1 :	CAACAGCAAATGATTGCCAGACACAC	(SEQ ID NO 6)
MIL-ICG-11 :	GAGGGGTTCCCGTCTGTAGTG	(SEQ ID NO 7)
15 MIL-ICG-22 :	TGAGGGGTTCTCGTCTGTAGTG	(SEQ ID NO 8)
MAC-ICG-1 :	CACTCGGTCGATCCGTGTGGA	(SEQ ID NO 9)
MAV-ICG-1 :	TCGGTCCGTCCGTGTGGAGTC	(SEQ ID NO 10)
20 MAV-ICG-22 :	GTGGCCGGCGTTCATCGAAA	(SEQ ID NO 11)
MIN-ICG-1 :	GCATAGTCCTTAGGGCTGATGCGTT	(SEQ ID NO 12)
MIN-ICG-2 :	GCTGATGCGTTCGTCGAAATGTGTA	(SEQ ID NO 13)
25 MIN-ICG-22 :	CTGATGCGTTCGTCGAAATGTGT	(SEQ ID NO 14)
MIN-ICG-222 :	TGATGCGTTCGTCGAAATGTGT	(SEQ ID NO 15)
MIN-ICG-2222 :	GGCTGATGCGTTCGTCGAAATGTGTA	(SEQ ID NO 16)
30 MAL-ICG-1 :	ACTAGATGAACCGCGTAGTCCTTGT	(SEQ ID NO 17)
MHEF-ICG-1 :	TGGACGAAAACCGGGTGCACAA	(SEQ ID NO 18)
MAH-ICG-1 :	GTGTAATTCTTTTAACTCTTGTGTAAAGTAAGTG	

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		(SEQ ID NO 19)
5	MCO-ICG-11 : TGGCCGGCGTGTTCATCGAAA	(SEQ ID NO 20)
	MTH-ICG-11 : GCACTTCAATTGGTGAAGTGCAGGCC	(SEQ ID NO 21)
	MTH-ICG-2 : GCGTGGTCTTCATGGCCGG	(SEQ ID NO 22)
	MEF-ICG-11 : ACGCGTGGTCCTCGTGG	(SEQ ID NO 23)
10	MSC-ICG-1 : TCGGCTCGTTCTGAGTGGTGTC	(SEQ ID NO 24)
	MKA-ICG-1 : GATGCGTTGCTACGGGTAGCGT	(SEQ ID NO 25)
	MKA-ICG-2 : GATGCGTTGCCTACGGGTAGCGT	(SEQ ID NO 26)
15	MKA-ICG-3 : ATGCGTTGCCCTACGGGTAGCGT	(SEQ ID NO 27)
	MKA-ICG-4 : CGGGCTCTGTTCGAGAGTTGTC	(SEQ ID NO 28)
20	MKA-ICG-5 : CCCTCAGGGATTTCTGGGTGTTG	(SEQ ID NO 182)
	MKA-ICG-6 : GGACTCGTCCAAGAGTGTGTC	(SEQ ID NO 183)
	MKA-ICG-7 : TCAGGGCTTGGCCAGAGCTGTT	(SEQ ID NO 184)
25	MKA-ICG-8 : GGGTGCGAACAGCAAGCGA	(SEQ ID NO 185)
	MKA-ICG-9 : GATGCGTTGCCCTACGGG	(SEQ ID NO 186)
	MKA-ICG-10 : CCCTACGGGTAGCGTGTCTTTG	(SEQ ID NO 187)
30	MCH-ICG-1 : GGTGTGGACTTGACTTCTGAATAG	(SEQ ID NO 29)
	MCH-ICG-2 : CGGCAAAACGTGGACTGTCA	(SEQ ID NO 30)
	MGO-ICG-1 : AACACCCTGGGTGCTGTCC	(SEQ ID NO 31)
35	MGO-ICG-2 : GTATGCGTTGTCGTTCGCGGC	(SEQ ID NO 32)
	MGO-ICG-5 : CGTGAGGGGTACCGTCTGTAG	(SEQ ID NO 33)
	MUL-ICG-1 : GGTTTGGGATGTTGTCACC	(SEQ ID NO 175)
40	MGV-ICG-1 : CGACTGAGGTGACGTGGTGT	(SEQ ID NO 176)
	MGV-ICG-2 : GGTGTTGAGCATTGAATAGTGGTGC	(SEQ ID NO 177)
	MXE-ICG-1 : GTTGGGCAGCAGGCAGTAACC	(SEQ ID NO 178)
45	MSI-ICG-1 : CCGGCAACGGTTACGTGTT	(SEQ ID NO 179)
	MFO-ICG-1 : TCGTTGGATGGCCTCGCACCT	(SEQ ID NO 180)
	MFO-ICG-2 : ACTTGGCGTGGGATGCGGGAA	(SEQ ID NO 181)
50	MML-ICG-1 : CGGATCGATTGAGTGCTTGTCCC	(SEQ ID NO 188)
	MML-ICG-2 : TCTAAATGAACGCCACTGCCGATGG	(SEQ ID NO 189)
	MCE-ICG-1 : TGAGGGAGCCCGTGCCTGTA	(SEQ ID NO 190)
55	MHP-ICG-1 : CATGTTGGCTTGATCGGGTGC	(SEQ ID NO 191)

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and more preferably to at least one probe of the following restricted group of spacer probes:

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	MYC-ICG-1 :	ACTGGATAGGGTGCAGCATCTA	(SEQ ID NO 1)
	MYC-ICG-22 :	CTTCTGAATAGGGTGCAGCATCT	(SEQ ID NO 2)
5	MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGCCA	(SEQ ID NO 3)
	MTB-ICG-2 :	GACTTGTCCAGGTGTTGCCAC	(SEQ ID NO 4)
	MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)
10	MAI-ICG-1 :	CAACAGCAAATGATTGCCAGACACAC	(SEQ ID NO 6)
	MIL-ICG-11 :	GAGGGGTTCCCGTCTGTAGTG	(SEQ ID NO 7)
	MIL-ICG-22 :	TGAGGGGTTCTCGTCTGTAGTG	(SEQ ID NO 8)
15	MAC-ICG-1 :	CACTCGGTCGATCCGTGTGGA	(SEQ ID NO 9)
	MAV-ICG-1 :	TCGGTCCCGTCCGTGTGGAGTC	(SEQ ID NO 10)
	MAV-ICG-22 :	GTGGCCGGCGTTCATCGAAA	(SEQ ID NO 11)
20	MIN-ICG-1 :	GCATAGTCCTAGGGCTGATGCGTT	(SEQ ID NO 12)
	MAL-ICG-1 :	ACTAGATGAACCGCTAGCCTTGT	(SEQ ID NO 17)
	MCO-ICG-11 :	TGGCCGGCGTGTTCATCGAAA	(SEQ ID NO 20)
25	MTH-ICG-11 :	GCACTTCAATTGGTGAAGTGCAGGCC	(SEQ ID NO 21)
	MTH-ICG-2 :	GCGTGGTCTTCATGGCCGG	(SEQ ID NO 22)
	MEF-ICG-11 :	ACGCGTGGTCCTCGTGG	(SEQ ID NO 23)
30	MSC-ICG-1 :	TCGGCTCGTTCTGAGTGGTGT	(SEQ ID NO 24)
	MKA-ICG-3 :	ATGCGTTGCCCTACGGGTAGCGT	(SEQ ID NO 27)
	MKA-ICG-4 :	CGGGCTCTGTCGAGAGTTGT	(SEQ ID NO 28)
35	MKA-ICG-5 :	CCCTCAGGGATTTCTGGGTGTTG	(SEQ ID NO 182)
	MKA-ICG-6 :	GGACTCGTCCAAGAGTGTGTC	(SEQ ID NO 183)
	MKA-ICG-7 :	TCGGGCTTGGCCAGAGCTGTT	(SEQ ID NO 184)
40	MKA-ICG-8 :	GGGTGCGAACAGCAAGCGA	(SEQ ID NO 185)
	MKA-ICG-9 :	GATGCGTTGCCCTACGGG	(SEQ ID NO 186)
	MKA-ICG-10 :	CCCTACGGGTAGCGTGTCTTTG	(SEQ ID NO 187)
45	MCH-ICG-1 :	GGTGTGGACTTGACTCTGAATAG	(SEQ ID NO 29)
	MCH-ICG-2 :	CGGCAAAACGTCGGACTGTCA	(SEQ ID NO 30)
	MCH-ICG-3 :	GGTGTGGCCTGACTTATGGATAG	(SEQ ID NO 210)
50	MGO-ICG-5 :	CGTGAGGGTCATCGTGTAG	(SEQ ID NO 33)
	MUL-ICG-1 :	GGTTTCGGGATGTTGTCCCACC	(SEQ ID NO 175)
	MGV-ICG-1 :	CGACTGAGGTCGACGTGGTGT	(SEQ ID NO 176)
55	MGV-ICG-2 :	GGTGTGGAGCATTGAATAGTGGTGC	(SEQ ID NO 177)

MGV-ICG-3 :	TCGGGCCCGCGTGTCTGTCAAA	(SEQ ID NO 211)
MXE-ICG-1 :	GTTGGGCAGCAGGCAGTAACC	(SEQ ID NO 178)
MSI-ICG-1 :	CCGGCAACGGTTACGTGTT	(SEQ ID NO 179)
MFO-ICG-1 :	TCGTTGGATGGCCTCGCACCT	(SEQ ID NO 180)
MFO-ICG-2 :	ACTTGGCGTGGGATGCAGGAA	(SEQ ID NO 181)
MML-ICG-1 :	CGGATCGATTGAGTGCTTGTCCC	(SEQ ID NO 188)
MML-ICG-2 :	TCTAAATGAACGCAGTGCCTGG	(SEQ ID NO 189)
MCE-ICG-1 :	TGAGGGAGCCCGTGCCTGTA	(SEQ ID NO 190)
MHP-ICG-1 :	CATGTTGGGCTTGATCGGGTGC	(SEQ ID NO 191)

or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 76-110, or 157-174 provided said probe hybridizes specifically to a Mycobacterium species.

[0126] The sequences represented by SEQ ID NO 76-110 and 157-174 are new.

[0127] Preferentially, at least two, three, four, five, six, seven, eight or more of said probes are used simultaneously.

[0128] As described above, the preferred restricted set of probes are those probes which showed a sensitivity and specificity of more than 80%, preferably more than 90%, most preferably more than 95%, under the specific hybridization conditions as described in the examples section.

[0129] In one specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium tuberculosis complex strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGGCCA	(SEQ ID NO 3)
MTB-ICG-2 :	GACTTGTCCAGGTGTTGTCCCAC	(SEQ ID NO 4)
MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)

or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 76 provided said probe hybridizes specifically to the M. tuberculosis complex. The M. tuberculosis complex comprises M. tuberculosis, M. bovis, M. bovis BCG, M. africanum and M. microti strains.

[0130] The sequence represented in SEQ ID NO 76 is new.

[0131] Preferentially, at least two, or three of said probes are used simultaneously.

[0132] In another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium strains from the MAIS-complex, wherein step (iii) comprises hybridizing to at least one of the following probes:

MAI-ICG-1 :	CAACAGCAAATGATTGCCAGACACAC	(SEQ ID NO 6)
MIL-ICG-11 :	GAGGGGTTCCCGTCTGTAGTG	(SEQ ID NO 7)
MIL-ICG-22 :	TGAGGGGTTCTCGTCTGTAGTG	(SEQ ID NO 8)
MAC-ICG-1 :	CACTCGGTCGATCCGTGTGGA	(SEQ ID NO 9)
MAV-ICG-1 :	TCGGTCCGTCCGTGTGGAGTC	(SEQ ID NO 10)
MAV-ICG-22 :	GTGGCCGGCGTTCATCGAAA	(SEQ ID NO 11)
MIN-ICG-1 :	GCATAGTCCTTAGGGCTGATGCGTT	(SEQ ID NO 12)
MIN-ICG-2 :	GCTGATGCGTTCGTCGAAATGTGTA	(SEQ ID NO 13)
MIN-ICG-22 :	CTGATGCGTTCGTCGAAATGTGT	(SEQ ID NO 14)
MIN-ICG-222 :	TGATGCGTTCGTCGAAATGTGT	(SEQ ID NO 15)
MIN-ICG-2222 :	GGCTGATGCGTTCGTCGAAATGTGTA	(SEQ ID NO 16)
MAL-ICG-1 :	ACTAGATGAACCGCGTAGTCCTTGT	(SEQ ID NO 17)
MHEF-ICG-1 :	TGGACGAAAACCGGGTGCACAA	(SEQ ID NO 18)
MAH-ICG-1 :	GTGTAATTCTTTAACTCTTGTGTAAAGTAAGTG	
		(SEQ ID NO 19)
MCO-ICG-11 :	TGGCCGGCGTGTTCATCGAAA	(SEQ ID NO 20)
MTH-ICG-11 :	GCACTTCAATTGGTGAAGTGCAGGCC	(SEQ ID NO 21)
MTH-ICG-2 :	GCGTGGTCTTCATGCCGG	(SEQ ID NO 22)
MEF-ICG-11 :	ACGCGTGGTCCTCGTGG	(SEQ ID NO 23)
MSC-ICG-1 :	TCGGCTCGTTCTGAGTGGTGTC	(SEQ ID NO 24)

or to equivalents of said probes,
and/or to any probe derived from SEQ ID NO 77-100 or 108-110, provided said probe hybridizes specifically to strains
from the MAIS complex. The MAIS complex as defined in this invention comprises all strains of M. avium, M. intracellulare and M. scrofulaceum and all strains closely related to the above mentioned species and not clearly belonging to another defined Mycobacterium species. The latter group of strains are defined in this invention as "MIC strains" (M. intracellulare complex).

[0133] Preferentially, at least two, three, four, five, six, seven, eight or more of said probes are used simultaneously.
[0134] In still another specific embodiment, the invention provides for a method as described above, to detect and identify one or more M. avium and M. paratuberculosis strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

MAV-ICG-1 : TCGGTCCGTCCGTGTGGAGTC (SEQ ID NO 10)

MAV-ICG-22 : GTGGCCGGCGTTCATCGAAA (SEQ ID NO 11)

or to equivalents of said probes,
and/or to any probe derived from SEQ ID NO 77 and 78 provided said probe hybridizes specifically to M. avium or M. paratuberculosis.

[0135] The sequences as represented in SEQ ID NO 77 and 78 are new.

[0136] Preferentially, this embodiment uses both probes in combination.

[0137] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium intracellulare strains and MIC-strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

MAI-ICG-1 : CAACAGCAAATGATTGCCAGACACAC (SEQ ID NO 6)

10 MIL-ICG-11 : GAGGGGTTCCCGTCTGTAGTG (SEQ ID NO 7)

MIL-ICG-22 : TGAGGGGTTCTCGTCTGTAGTG (SEQ ID NO 8)

MAC-ICG-1 : CACTCGGTGATCCGTGTGGA (SEQ ID NO 9)

15 MIN-ICG-1 : GCATAGTCCTTAGGGCTGATGCGTT (SEQ ID NO 12)

MIN-ICG-2 : GCTGATGCGTTCGTCGAAATGTGTA (SEQ ID NO 13)

MIN-ICG-22 : CTGATGCGTTCGTCGAAATGTGT (SEQ ID NO 14)

20 MIN-ICG-222 : TGATGCGTTCGTCGAAATGTGT (SEQ ID NO 15)

MIN-ICG-2222 : GGCTGATGCGTTCGTCGAAATGTGTA (SEQ ID NO 16)

25 MAL-ICG-1 : ACTAGATGAACGCGTAGTCCTTGT (SEQ ID NO 17)

MHEF-ICG-1 : TGGACGAAAACCAGGGTGCACAA (SEQ ID NO 18)

MAH-ICG-1 : GTGTAATTCTTTAACTCTGTGTAAAGTAAGTG (SEQ ID NO 19)

30 MCO-ICG-11 : TGGCCGGCGTGTTCATCGAAA (SEQ ID NO 20)

MTH-ICG-11 : GCACTTCAATTGGTGAAGTGCAGGCC (SEQ ID NO 21)

35 MTH-ICG-2 : GCGTGGTCTTCATGGCCGG (SEQ ID NO 22)

MEF-ICG-11 : ACGCGTGGTCCTCGTGG (SEQ ID NO 23)

or to equivalents of said probes,

40 and/or to any probe derived from SEQ ID NO 79, 80, 81, 82, 83, 84, 85, 86, 87, 88, 89, 90, 91, 92, 93, 94, 95, 96, 97, 98 or 99 provided said probe hybridizes specifically to M. intracellulare strains and MIC-strains.

[0138] The sequences as represented in SEQ ID NO 79, 80, 81, 82, 83, 84, 85, 86, 87, 88, 89, 90, 91, 92, 93, 94, 95, 96, 97, 98 or 99 are new.

[0139] Preferentially, at least two, three, four, five, six, seven, eight or more of said probes are used simultaneously.

45 [0140] In still another specific embodiment, the invention provides for a method as described above, to detect and identify one or more Mycobacterium intracellulare strains in a sample, wherein step (iii) comprises hybridizing to at least the following probes :

50 MIN-ICG-1 : GCATAGTCCTTAGGGCTGATGCGTT (SEQ ID NO 12)

or to equivalents of said probe,

and/or to any probe derived from SEQ ID NO 89 provided said probe hybridizes specifically to M. intracellulare strains.

55 [0141] In still another specific embodiment, the invention provides for a method as described above, to detect and identify one or more Mycobacterium scrofulaceum strains in a sample, wherein step (iii) comprises hybridizing to the following probe:

MSC-ICG-1 : TCGGCTCGTCTGAGTGGTGTC

(SEQ ID NO 24)

or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 100 provided said probe hybridizes specifically to M. scrofulaceum.

[0142] The sequence as represented in SEQ ID NO 100 is new.

[0143] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium kansasii strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

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MKA-ICG-1 : GATGCGTTGCTACGGTAGCGT (SEQ ID NO 25)

MKA-ICG-2 : GATGCGTTGCCTACGGTAGCGT (SEQ ID NO 26)

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MKA-ICG-3 : ATGCGTTGCCCTACGGTAGCGT (SEQ ID NO 27)

MKA-ICG-4 : CGGGCTCTGTTGAGAGTTGTC (SEQ ID NO 28)

20

MKA-ICG-5 : CCCTCAGGGATTCTGGGTGTT (SEQ ID NO 182)

MKA-ICG-6 : GGACTCGTCCAAGAGTGTGTC (SEQ ID NO 183)

MKA-ICG-7 : TCGGGCTTGGCCAGAGCTGTT (SEQ ID NO 184)

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MKA-ICG-8 : GGGTGCACACAGCAAGCGA (SEQ ID NO 185)

MKA-ICG-9 : GATGCGTTGCCCTACGGG (SEQ ID NO 186)

MKA-ICG-10 : CCCTACGGTAGCGTGTCTTTG (SEQ ID NO 187)

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and more preferably to :

MKA-ICG-3 : ATGCGTTGCCCTACGGTAGCGT (SEQ ID NO 27)

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MKA-ICG-4 : CGGGCTCTGTTGAGAGTTGTC (SEQ ID NO 28)

MKA-ICG-5 : CCCTCAGGGATTCTGGGTGTT (SEQ ID NO 182)

40

MKA-ICG-6 : GGACTCGTCCAAGAGTGTGTC (SEQ ID NO 183)

MKA-ICG-7 : TCGGGCTTGGCCAGAGCTGTT (SEQ ID NO 184)

MKA-ICG-8 : GGGTGCACACAGCAAGCGA (SEQ ID NO 185)

45

MKA-ICG-9 : GATGCGTTGCCCTACGGG (SEQ ID NO 186)

MKA-ICG-10 : CCCTACGGTAGCGTGTCTTTG (SEQ ID NO 187)

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or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 101, 167, 168 or 169 provided said probe hybridizes specifically to M. kansasii.

[0144] The sequences as represented in SEQ ID NO 101, 167, 168 and 169 are new.

[0145] Preferentially, at least two, three or four of said probes are used simultaneously.

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[0146] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium chelonae strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

MCH-ICG-1 :	GGTGTGGACTTGACTTCTGAATAG	(SEQ ID NO 29)
MCH-ICG-2 :	CGGCAAAACGTCGGACTGTCA	(SEQ ID NO 30)
MCH-ICG-3 :	GGTGTGGCCTTGACTTATGGATAG	(SEQ ID NO 210)

or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 102, 103 or 174 provided said probe hybridizes specifically to M. chelonae.

According to another preferential embodiment, these three probes are used in combination.

[0147] The sequences as represented in SEQ ID NO 102, 103 and 174 are new.

[0148] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium gordonaee strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

MGO-ICG-1 :	AACACCCTGGGTGCTGTCC	(SEQ ID NO 31)
MGO-ICG-2 :	GTATGCGTTGTCGTTCGCGGC	(SEQ ID NO 32)
MGO-ICG-5 :	CGTGAGGGGTCATCGTCTGTAG	(SEQ ID NO 33)

and more preferably to:

MGO-ICG-5 :	CGTGAGGGGTCATCGTCTGTAG	(SEQ ID NO 33)
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or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 104, 105 or 106 provided said probe hybridizes specifically to M. gordonaee.

[0149] The sequences as represented in SEQ ID NO 104 to 106 are new.

[0150] Preferentially, at least two or three of said probes are used simultaneously.

[0151] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium ulcerans strains or Mycobacterium marinum strains in a sample, wherein step (iii) comprises hybridizing to the following probe:

MUL-ICG-1 :	GGTTTCGGGATGTTGTCCCACC	(SEQ ID NO 175)
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or to equivalents of said probe,

and/or to any probe derived from SEQ ID NO 157 provided said probe hybridizes specifically to M. ulcerans and M. marinum.

[0152] The sequence as represented in SEQ ID NO 157 is new.

[0153] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium genavense strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

MGV-ICG-1 :	CGACTGAGGTGACGTGGTGT	(SEQ ID NO 176)
MGV-ICG-2 :	GGTGTGAGCATTGAATACTGGTTGC	(SEQ ID NO 177)
MGV-ICG-3 :	TCGGGCCGCGTGTGTCGTAAA	(SEQ ID NO 211)

or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 158, 159, 160, 161 or 162 provided said probe hybridizes specifically to M. genavense.

[0154] The sequences as represented in SEQ ID NO 158 to 162 are new.

[0155] As described in the examples, M. genavense includes M. genavense strains sensu strictu and a group of closely related strains called M. simiae-like. The former group of strains can be detected specifically with probe MGV-

ICG-1 while the latter group hybridizes specifically with probe MGV-ICG-3. Probe MGV-ICG-2 detects both groups.

[0156] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium xenopi strains in a sample, wherein step (iii) comprises hybridizing to the following probe:

5

MXE-ICG-1 : GTTGGGCAGCAGGCAGTAACC (SEQ ID NO 178)

10 or to equivalents of said probe,

and/or to any probe derived from SEQ ID NO 163 provided said probe hybridizes specifically to M. xenopi.

[0157] The sequence as represented in SEQ ID NO 163 is new.

[0158] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium simiae strains in a sample, wherein step (iii) comprises hybridizing to the following probe:

15

MSI-ICG-1 : CCGGCAACGGTTACGTGTTC (SEQ ID NO 179)

20 or to equivalents of said probe,

and/or to any probe derived from SEQ ID NO 164 or 165 provided said probe hybridizes specifically to M. simiae.

[0159] The sequence as represented in SEQ ID NO 164 or 165 is new.

[0160] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium fortuitum strains in a sample, wherein step (iii) comprises hybridizing to at least one of the the following probes:

25

MFO-ICG-1 : TCGTTGGATGGCCTCGCACCT (SEQ ID NO 180)

30

MFO-ICG-2 : ACTTGGCGTGGGATGCGGGAA (SEQ ID NO 181)

or to equivalents of said probes or to any probe derived from SEQ ID NO 166 provided said probe hybridizes specifically to M. fortuitum.

[0161] The sequence as represented in SEQ ID NO 166 is new.

[0162] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium celatum strains in a sample, wherein step (iii) comprises hybridizing to the following probe:

35

40

MCE-ICG-1 : TGAGGGAGCCCGTGCCTGTA (SEQ ID NO 190)

or to equivalents of said probe,

and/or to any probe derived from SEQ ID NO 170 provided said probe hybridizes specifically to M. celatum.

[0163] The sequence as represented in SEQ ID NO 170 is new.

[0164] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium haemophilum strains in a sample, wherein step (iii) comprises hybridizing to the following probe:

45

50

MHP-ICG-1 : CATGTTGGCTTGATCGGGTGC (SEQ ID NO 191)

or to equivalents of said probe,

and/or to any probe derived from SEQ ID NO 171, 172 or 173 provided said probe hybridizes specifically to M. haemophilum.

[0165] The sequences as represented in SEQ ID NO 171 to 173 are new.

[0166] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium malmoense strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

MML-ICG-1 : CGGATCGATTGAGTGCTTGTCCC (SEQ ID NO 188)
 MML-ICG-2 : TCTAAATGAACGCAGTGCCGATGG (SEQ ID NO 189)

5

or to equivalents of said probes,
 and/or to any probe derived from SEQ ID NO 107 provided said probe hybridizes specifically to M. malmoense.
 [0167] The sequence as represented in SEQ ID NO 107 is new.
 [0168] In still another specific embodiment, the invention provides for a method as described above to detect and identify one or more Mycobacterium strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

15

MYC-ICG-1 : ACTGGATAGTGGTTGCGAGCATCTA (SEQ ID NO 1)
 MYC-ICG-22 : CTTCTGAATAGTGGTTGCGAGCATCT (SEQ ID NO 2)

or to equivalents of said probes.
 [0169] According to a preferred embodiment, both probes are used in combination.
 [0170] The invention also provides for a method as described above to detect and identify one or more Mycoplasma strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

25

MPN-ICG 1 : ATCGGTGGTAAATTAAACCCAAATCCCTGT (SEQ ID NO 49)
 MPN-ICG 2 : CAGTTCTGAAAGAACATTCCGCTTCTTC (SEQ ID NO 50)
 MGE-ICG 1 : CACCCATTAATTTTCGGTGTAAAACCC (SEQ ID NO 51)
 Mycoplasma-ICG : CAAAAGTAAAAACGACAATCTTCTAGTTCC (SEQ ID NO 52)

30

or to equivalents of said probes,
 and/or to any probe derived from SEQ ID NO 124 or 125 provided said probe hybridizes specifically with Mycoplasma species.
 [0171] Preferentially, at least two, three or four of said probes are used simultaneously.
 [0172] More particularly, the invention provides for a method as described above to detect and identify one or more Mycoplasma pneumoniae strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

40

MPN-ICG 1 : ATCGGTGGTAAATTAAACCCAAATCCCTGT (SEQ ID NO 49)
 MPN-ICG 2 : CAGTTCTGAAAGAACATTCCGCTTCTTC (SEQ ID NO 50)

45

or to equivalents of said probes,
 and/or to any probe derived from SEQ ID NO 125 provided said probe hybridizes specifically to Mycoplasma pneumoniae. According to a preferred embodiment, both these probes are used in combination.
 [0173] The sequence as represented in SEQ ID NO 125 is new.
 [0174] In another particular embodiment, the invention provides for a method as described above to detect and identify one or more Mycoplasma genitalium strains in a sample, wherein step (iii) comprises hybridizing to the following probe:

MGE-ICG 1 : CACCCATTAATTTTCGGTGTAAAACCC (SEQ ID NO 51)

55

or to equivalents of said probes,
 and/or to any probe derived from SEQ ID NO 124 provided said probe hybridizes specifically to Mycoplasma genitalium.
 [0175] The sequence as represented in SEQ ID NO 124 is new.

[0176] The invention also provides for a method as described above to detect and identify one or more Pseudomonas strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

5	PA-ICG 1 : TGGTGTGCTGCGTGATCCGAT	(SEQ ID NO 34)
	PA-ICG 2 : TGAATGTTCTGGATGAACATTGATT	(SEQ ID NO 35)
	PA-ICG 3 : CACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 36)
10	PA-ICG 4 : TGAATGTTCTGT(G/A)(G/A)ATGAACATTGATTCTGGTC	(SEQ ID NO 37)
	PA-ICG 5 : CTCTTCACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 38)

15 or to equivalents of said probes,
and/or to any probe derived from SEQ ID NO 111, 112, 113, 114 or 115 provided said probe hybridizes specifically to Pseudomonas strains.

- [0177] The sequences as represented in SEQ ID NO 111 to 115 are new.
 20 [0178] Preferentially, at least two, three or four of said probes are used simultaneously.
 [0179] More particularly, the invention provides for a method as described above to detect and identify one or more Pseudomonas aeruginosa strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

25	PA-ICG 1 : TGGTGTGCTGCGTGATCCGAT	(SEQ ID NO 34)
	PA-ICG 2 : TGAATGTTCTGGATGAACATTGATT	(SEQ ID NO 35)
	PA-ICG 3 : CACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 36)
30	PA-ICG 4 : TGAATGTTCTGT(G/A)(G/A)ATGAACATTGATTCTGGTC	(SEQ ID NO 37)
	PA-ICG 5 : CTCTTCACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 38)

35 and most preferably to at least one of the following probes:

40	PA-ICG 1 : TGGTGTGCTGCGTGATCCGAT	(SEQ ID NO 34)
	PA-ICG 4 : TGAATGTTCTGT(G/A)(G/A)ATGAACATTGATTCTGGTC	(SEQ ID NO 37)
	PA-ICG 5 : CTCTTCACTGGTGATCATTCAAGTCAAG	(SEQ ID NO 38)

45 or to equivalents of said probes,
and/or to any probe derived from SEQ ID NO 111 provided said probe hybridizes specifically to Pseudomonas aeruginosa.

- [0180] The sequence as represented in SEQ ID NO 111 is new.
 50 [0181] Preferentially, at least two, three, four or five of said probes are used simultaneously.
 [0182] The invention also provides for a method as described above to detect and identify one or more Staphylococcus species in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

	STAU-ICG 1 : TACCAAGCAAAACCGAGTGAATAAAGAGTT	(SEQ ID NO 53)
	STAU-ICG 2 : CAGAACGATGCGGAATAACGTGAC	(SEQ ID NO 54)
5	STAU-ICG 3 : AACGAAGCCGTATGTGAGCATTGAC	(SEQ ID NO 55)
	STAU-ICG 4 : GAACGTAACCTCATGTTAACGTTGACTTAT	(SEQ ID NO 56)

or to equivalents of said probes,
 10 and/or to any probe derived from SEQ ID NO 139, 140, 141, 142, 143 or 144 provided said probe hybridizes specifically to Staphylococcus species.

[0183] The sequences as represented in SEQ ID NO 139 to 144 are new.
 [0184] Preferentially, at least two, three or four of said probes are used simultaneously.
 [0185] More particularly, the invention provides for a method as described above to detect and identify one or more Staphylococcus aureus strains in a sample, wherein step (iii) comprises hybridizing to at least one, and preferably both of the following probes:

	STAU-ICG 3 : AACGAAGCCGTATGTGAGCATTGAC	(SEQ ID NO 55)
20	STAU-ICG 4 : GAACGTAACCTCATGTTAACGTTGACTTAT	(SEQ ID NO 56)

or to equivalent of said probes,
 25 and/or to any probe derived from SEQ ID NO 139, 140, 141, 142, or 143 provided said probe hybridizes specifically to Staphylococcus aureus. According to a preferred embodiment, both these probes are used in combination.

[0186] In another specific embodiment the invention provides for a method as described above to detect and identify one or more Staphylococcus epidermidis strains in a sample, wherein step (iii) comprises hybridizing to any probe derived from SEQ ID NO 144 as long as this probe can be caused to hybridize specifically to Staphylococcus epidermidis.
 30 [0187] The invention also provides for a method as described above to detect and identify one or more Acinetobacter strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

	ACI-ICG 1 : GCTTAAGTGCACAGTGCTCTAAACTGA	(SEQ ID NO 57)
35	ACI-ICG 2 : CACGGTAATTAGTGTGATCTGACGAAG	(SEQ ID NO 58)

or to equivalents of said probes,
 40 and/or to any probe derived from SEQ ID NO 126, 127, 128, 129 or 130 provided said probe hybridizes specifically to Acinetobacter sp.. According to a preferred embodiment, both these probes are used in combination.
 [0188] The sequences as represented in SEQ ID NO 126 to 130 are new.
 [0189] More particularly, the invention provides for a method as described above to detect and identify one or more Acinetobacter baumanii strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:
 45

	ACI-ICG 1 : GCTTAAGTGCACAGTGCTCTAAACTGA	(SEQ ID NO 57)
50	ACI-ICG 2 : CACGGTAATTAGTGTGATCTGACGAAG	(SEQ ID NO 58)

or to equivalents of said probes,
 and/or to any probe derived from SEQ ID NO 126 provided said probe hybridizes specifically to Acinetobacter baumanii. According to a preferred embodiment, both these probes are used in combination.
 55 [0190] The invention also provides for a method as described above, to detect and identify one or more Listeria strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

LIS-ICG 1 : CAAGTAACCGAGAATCATCTGAAAGTGAATC (SEQ ID NO 39)

LMO-ICG 1 : AAACAACCTTACTTCGTAGAAGTAAATTGGTTAAG

5 (SEQ ID NO 40)

LMO-ICG 2 : TGAGAGGTTAGTACTTCTCAGTATGTTGTT (SEQ ID NO 41)

LMO-ICG 3 : AGGCACATGCTGAAGCATCGC (SEQ ID NO 42)

10 LIV-ICG 1 : GTTAGCATAAAATAGGTAACATTATGACACAGTAAC

(SEQ ID NO 43)

LSE-ICG 1 : AGTTAGCATAAGTAGTGTAACTATTATGACACAGAAG

15 LIS-ICG 1 : CGTTTCATAAGCGATCGCACGTT (SEQ ID NO 212)

and most preferably to at least one of the following probes:

20 LIS-ICG 1 : CAAGTAACCGAGAATCATCTGAAAGTGAATC (SEQ ID NO 39)

LMO-ICG 3 : AGGCACATGCTGAAGCATCGC (SEQ ID NO 42)

25 LIS-ICG 1 : CGTTTCATAAGCGATCGCACGTT (SEQ ID NO 212)

or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 116, 118, 119, 120, 121, 213, 214 or 215 provided said probe hybridizes specifically to Listeria species.

30 [0191] As described in the examples section, Listeria species encompass Listeria species *sensu strictu*, and a group of closely related organisms referred to as "Listeria-like organisms". The latter group can be specifically recognized by probe LIS-ICG 1.

[0192] The sequences as represented in SEQ ID NO 116, 118 to 121 and 213 to 215 are new.

[0193] Preferentially, at least two, three, four, five or six of said probes are used simultaneously.

35 [0194] More particularly, the invention provides for a method as described above, to detect and identify one or more Listeria monocytogenes strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

40 LMO-ICG 1 : AAACAACCTTACTTCGTAGAAGTAAATTGGTTAAG (SEQ ID NO 40)

LMO-ICG 2 : TGAGAGGTTAGTACTTCTCAGTATGTTGTT (SEQ ID NO 41)

LMO-ICG 3 : AGGCACATGCTGAAGCATCGC (SEQ ID NO 42)

45 and most preferably to the following probe:

LMO-ICG 3 : AGGCACATGCTGAAGCATCGC (SEQ ID NO 42)

50 or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 120 provided said probe hybridizes specifically to Listeria monocytogenes.

[0195] Preferentially, at least two, or three of said probes are used simultaneously.

55 [0196] The invention also provides for a method as described above to detect and identify one or more Brucella strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

BRU-ICG 1 :	CGTGCCGCCTCGTTCTCTTT	(SEQ ID NO 59)
BRU-ICG 2 :	TTCGCTTCGGGGTGGATCTGTG	(SEQ ID NO 60)
5 BRU-ICG 3 :	GCGTAGTAGCGTTGCGTCGG	(SEQ ID NO 193)
BRU-ICG 4 :	CGCAAGAAGCTTGCTCAAGCC	(SEQ ID NO 194)

and most preferably to at least one of the following probes:

10	BRU-ICG 2 :	TTCGCTTCGGGGTGGATCTGTG	(SEQ ID NO 60)
15	BRU-ICG 3 :	GCGTAGTAGCGTTGCGTCGG	(SEQ ID NO 193)
	BRU-ICG 4 :	CGCAAGAAGCTTGCTCAAGCC	(SEQ ID NO 194)

or to equivalents of said probes,

- 20 and/or to any probe derived from SEQ ID NO 131, 132 or 154 provided said probe hybridizes specifically to Brucella strains.
[0197] The sequences as represented in SEQ ID NO 131, 132 and 154 are new.
[0198] The invention also provides for a method as described above to detect and identify one or more Salmonella strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

25	SALM-ICG 1 :	CAAAACTGACTTACGAGTCACGTTGAG	(SEQ ID NO 61)
	SALM-ICG 2 :	GATGTATGCTTCGTTATTCCACGCC	(SEQ ID NO 62)
30	STY-ICG 1 :	GGTCAAACCTCCAGGGACGCC	(SEQ ID NO 63)
	SED-ICG 1 :	GCGGTAATGTGTGAAAGCGTTGCC	(SEQ ID NO 64)

and most preferably to the following probe:

35	SALM-ICG 1 :	CAAAACTGACTTACGAGTCACGTTGAG	(SEQ ID NO 61)
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or to equivalents of said probes,

- 40 and/or to any probe derived from SEQ ID NO 133, 134, 135, 136, 137 or 138 provided said probe hybridizes specifically to Salmonella strains.
[0199] The sequences as represented in SEQ ID NO 133 to 138 are new.
[0200] Preferentially, at least two, three, or four of said probes are used simultaneously.
[0201] The invention also relates to a method as described above to detect and identify one or more Chlamydia strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

45	CHTR-ICG 1 :	GGAAGAACGCTGAGAAGGTTCTGAC	(SEQ ID NO 45)
50	CHTR-ICG 2 :	GCATTATATGTAAGAGCAAGCATTCTATTCA	(SEQ ID NO 46)
	CHTR-ICG 3 :	GAGTAGCGTGGTGAGGACGAGA	(SEQ ID NO 47)
	CHTR-ICG 4 :	GAGTAGCGCGGTGAGGACGAGA	(SEQ ID NO 201)
55	CHPS-ICG 1 :	GGATAACTGTCTAGGACGGTTGAC	(SEQ ID NO 48)

or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 122, 123 or 197 provided that said probe hybridizes specifically to Chlamy-

dia strains.

[0202] Preferentially, at least two, three, four or five of said probes are used simultaneously.

[0203] More particularly, the invention relates to a method as described above to detect and identify one or more Chlamydia trachomatis strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes:

CHTR-ICG 1 :	GGAAGAACCTGAGAAGGTTCTGAC	(SEQ ID NO 45)
10 CHTR-ICG 2 :	GCATTATATGTAAGAGCAAGCATTCTATTCA	(SEQ ID NO 46)
CHTR-ICG 3 :	GAGTAGCGTGGTGAGGACGAGA	(SEQ ID NO 47)
CHTR-ICG 4 :	GAGTAGCGCGGTGAGGACGAGA	(SEQ ID NO 201)

15 or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 123 or 197 provided said probe hybridizes specifically to Chlamydia trachomatis.

[0204] The sequences as represented in SEQ ID NO 123 and 197 are new.

20 [0205] Preferentially, at least two, three or four of said probes are used simultaneously.

[0206] In another particular embodiment, the invention provides for a method as described above to detect and identify one or more Chlamydia psittaci strains in a sample, wherein step (iii) comprises hybridizing to at least the following probe:

CHPS-ICG 1 :	GGATAACTGTCTAGGACGGTTGAC	(SEQ ID NO 48)
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or to equivalents of said probe,

and/or to any probe derived from SEQ ID NO 122 provided said probe hybridizes specifically to Chlamydia psittaci.

30 [0207] The sequence of SEQ ID NO 122 is new.

[0208] The invention also provides for a method as described above, to detect one or more Streptococcus strains in a sample, wherein step (iii) comprises hybridizing to any probe derived from SEQ ID NO 145, 146, 147, 148, 149, 150, 151, 152 or 153 provided said probe hybridizes specifically to Streptococcus strains, or equivalents of these probes.

35 [0209] The sequences as represented in SEQ ID NO 145, 146, 147, 148, 149, 150, 151, 152 or 153 are new.

[0210] The invention also provides for a method as described above, to detect one or more Yersinia enterocolitica strains in a sample, wherein step (iii) comprises hybridizing to at least one of the following probes :

YEC-ICG 1 :	GGAAAAGGTACTGCACGTGACTG	(SEQ ID NO 198)
YEC-ICG 2 :	GACAGCTGAAACTTATCCCTCCG	(SEQ ID NO 199)
YEC-ICG 3 :	GCTACCTGTTGATGTAATGAGTCAC	(SEQ ID NO 200)

45 or to equivalents of said probes,

and/or to any probe derived from SEQ ID NO 195 or 196, provided said probe hybridizes specifically to Yersinia enterocolitica.

[0211] The sequences as represented in SEQ ID NO 195 and 196 are new.

50 [0212] In some cases it may be advantageous to amplify not all organisms present in a sample, but only more specific taxa, which are considered to be relevant. In these cases the invention provides for primers allowing the specific amplification of the spacer region for only those beforehand defined taxa.

[0213] The invention thus provides for a method as described above to detect and identify specifically Chlamydia trachomatis in a sample, wherein step (ii) comprises amplification of the 16S-23S rRNA spacer region or a part of it, using at least one of the following primers:

CHTR-P1 : AAGGTTCTGACTAGGTTGGGC (SEQ ID NO 69)

CHTR-P2 : GGTGAAGTGCTTGCATGGATCT (SEQ ID NO 70)

5

or equivalents of these primers, said equivalents differing in sequence from the above mentioned primers by changing one or more nucleotides, provided that said equivalents still amplify specifically the spacer region or part of it from Chlamydia trachomatis.

[0214] Preferably both primers are used.

10 [0215] The invention also provides for a method as described above to detect and identify specifically Listeria species in a sample, wherein step (ii) comprises amplification of the 16S-23S rRNA spacer region or a part of it, using at least one of the following primers:

15

LIS-P1 : ACCTGTGAGTTTCGTTCTCTC (SEQ ID NO 71)

LIS-P2 : CTATTGTTCACTTTGAGAGGTT (SEQ ID NO 72)

LIS-P3 : ATTTCCGTATCAGCGATGATAC (SEQ ID NO 73)

20

LIS-P4 : ACGAAGTAAAGGTTGTTTCT (SEQ ID NO 74)

LIS-P5 : GAGAGGTTACTCTCTTTATGTCAG (SEQ ID NO 75)

25

LIS-P6 : CTTTATGTCAGATAAAGTATGCAA (SEQ ID NO 202)

LIS-P7 : CGTAAAAGGGTATGATTATTG (SEQ ID NO 203)

or equivalents of these primers, said equivalents differing in sequence from the above mentioned primers by changing one or more nucleotides, provided that said equivalents still amplify specifically the spacer region or part of it from Listeria species.

30 [0216] The invention also relates to a method as described above to detect and identify specifically Mycobacterium species in a sample, wherein step (ii) comprises amplification of the 16S-23S rRNA spacer region or a part of it, using at least one of the following primers:

35

MYC-P1: TCCCTTGTGGCCTGTGTG (SEQ ID NO 65)

MYC-P2: TCCTTCATCGGCTCTCGA (SEQ ID NO 66)

40

MYC-P3: GATGCCAAGGCATCCACC (SEQ ID NO 67)

MYC-P4: CCTCCCACGTCTTCATCG (SEQ ID NO 68)

45

MYC-P5: CCTGGGTTGACATGCACAG (SEQ ID NO 192)

or equivalents of these primers, said equivalents differing in sequence from the above mentioned primers by changing one or more nucleotides, provided that said equivalents still amplify specifically the spacer region or part of it from Mycobacterium species.

50 [0217] The invention also provides for a method as described above to detect and identify specifically Brucella species in a sample, wherein step (ii) comprises amplification of the 16S-23S rRNA spacer region or part of it, using at least one of the following primers :

55

BRU-P1 :	TCGAGAATTGGAAAGAGGTC	(SEQ ID NO 204)
BRU-P2 :	AAGAGGTCGGATTATCCG	(SEQ ID NO 205)
5 BRU-P3 :	TTCGACTGCAAATGCTCG	(SEQ ID NO 206)
BRU-P4 :	TCTTAAAGCCGCATTATGC	(SEQ ID NO 207)

or equivalents of these primers, said equivalents differing in sequence from the above-mentioned primers by changing one or more nucleotides, provided that said equivalents still amplify specifically the spacer region of part of it from Brucella species.

10 [0218] The invention also provides for a method as described above to detect and identify specifically Yersinia enterocolitica species in a sample, wherein step (ii) comprises amplification of the 16S-23S rRNA spacer region or part of it, using at least one of the following primers :

15

YEC-P1 :	CCTAATGATATTGATTGCGC	(SEQ ID NO 208)
YEC-P2 :	ATGACAGGTTAACCTTACCCC	(SEQ ID NO 209)

20

or equivalents of these primers, said equivalents differing in sequence from the above-mentioned primers by changing one or more nucleotides, provided that said equivalents still amplify specifically the spacer region of part of it from Yersinia enterocolitica species.

25 [0219] The invention also provides for a composition comprising at least one of the probes and/or primers as defined above.

[0220] Said composition may comprise any carrier, support, label or diluent known in the art for probes or primers, more particularly any of the labels or supports detailed in the definitions section.

[0221] The invention relates more particularly to isolated probes and primers as defined above, more particularly any of the probes as specified in Table 1a or any of the primers as specified in Table 1b.

30 [0222] According to another embodiment, the present invention relates also to new spacer region sequences as defined above and as set out in figures 1-103 (SEQ ID NO 76 to 154, SEQ ID NO 157 to 174, SEQ ID NO 195 to 197 and SEQ ID NO 213 to 215).

[0223] In another embodiment the invention provides for a reverse hybridization method comprising any of the probes as defined above, wherein said probes are immobilized on a known location on a solid support, more preferably on a membrane strip.

35 [0224] In yet another embodiment the invention provides for a kit for the detection and identification of at least one micro-organism, or the simultaneous detection and identification of several micro-organisms in a sample, comprising the following components:

- 40 (i) when appropriate, at least one suitable primer pair to allow amplification of the intercistronic 16S-23S rRNA spacer region, or a part of it;
- (ii) at least one of the probes as defined above;
- (iii) a buffer, or components necessary to produce the buffer, enabling a hybridization reaction between said probes and the polynucleic acids present in the sample, or the amplified products thereof;
- 45 (iv) a solution, or components necessary to produce the solution, enabling washing of the hybrids formed under the appropriate wash conditions;
- (v) when appropriate, a means for detecting the hybrids resulting from the preceding hybridization.

FIGURE LEGENDS

50

[0225]

- Fig 1 : represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium tuberculosis strain H37RV ATCC 27294 (SEQ ID NO 76)
- 55 Fig 2: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium avium ATCC 151.769 (ITG 4991) (SEQ ID NO 77)

- Fig 3: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium paratuberculosis strains 316F and 2E (SEQ ID NO 78)
- 5 Fig 4: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 5513 (SEQ ID NO 79)
- Fig 5: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8695 (SEQ ID NO 80)
- 10 Fig 6: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8708 (SEQ ID NO 81).
- Fig 7: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8715 (SEQ ID NO 82)
- 15 Fig 8: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8054 (SEQ ID NO 83)
- Fig 9: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8737 (SEQ ID NO 84)
- 20 Fig 10: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8743 (SEQ ID NO 85)
- Fig 11: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8745 (SEQ ID NO 86)
- 25 Fig 12: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8748 (SEQ ID NO 87)
- Fig 13: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 8752 (SEQ ID NO 88)
- 30 Fig 14: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium intracellulare serovar 12 ITG 5915 (SEQ ID NO 89)
- Fig 15: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium lufu ITG 4755 (SEQ ID NO 90)
- 40 Fig 16: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 5922 (SEQ ID NO 91)
- Fig 17: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 1329 (SEQ ID NO 92)
- 45 Fig 18: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 1812 (SEQ ID NO 93)
- Fig 19: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 5280 (SEQ ID NO 94)
- 50 Fig 20: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 5620 (SEQ ID NO 95)
- Fig 21: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium strain ITG 5765 (SEQ ID NO 96)
- 55 Fig 22: represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium ITG 7395 (SEQ

ID NO 97)

Fig 23 : represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium ITG 8738 (SEQ ID NO 98)

5 Fig 24 : represents the DNA sequence of the 16S-23S rRNA spacer region from Mycobacterium ITG 926 (SEQ ID NO 99)

10 Fig 25 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium scrofulaceum ITG 4988 (SEQ ID NO 100)

Fig 26 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium kansasii ATCC 22478 (=ITG 4987) (SEQ ID NO 101)

15 Fig 27 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium chelonae abcessus ITG 4975 (SEQ ID NO 102)

20 Fig 28 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium chelonae chelonae ITG 9855 (SEQ ID NO 103)

25 Fig 29 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium gordonaee ITG 7703 (SEQ ID NO 104)

Fig 30 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium gordonaee ITG 7836 (SEQ ID NO 105)

30 Fig 31 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium gordonaee ITG 8059 (SEQ ID NO 106)

35 Fig 32 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium malmoense ITG 4842 and ITG 4832 (SEQ ID NO 107)

Fig 33 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium strain 8757 (SEQ ID NO 108)

40 Fig 34 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium ITG 8723 (SEQ ID NO 109)

Fig 35 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium ITG 8724 (SEQ ID NO 110)

45 Fig 36 : represents the DNA sequence of the 16S-23S spacer region from Pseudomonas aeruginosa UZG 5669 (SEQ ID NO 111)

Fig 37 : represents the DNA sequence of the 16S-23S spacer region from Pseudomonas pseudoalcaligenes LMG 1225 (SEQ ID NO 112)

50 Fig 38 : represents the DNA sequence of the 16S-23S spacer region from Pseudomonas stutzeri LMG 2333 (SEQ ID NO 113)

Fig 39 : represents the DNA sequence of the 16S-23S spacer region from Pseudomonas alcaligenes LMG 1224 (SEQ ID NO 114)

55 Fig 40 : represents the DNA sequence of the 16S-23S spacer region from Pseudomonas putida LMG 2232 (SEQ ID NO 115)

Fig 41 : represents the DNA sequence of the small 16S-23S spacer region from Listeria ivanovii CIP 7842 (SEQ ID NO 116)

- Fig 42 : represents the DNA sequence of the small 16S-23S spacer region from Listeria monocytogenes (SEQ ID NO 117)
- 5 Fig 43 : represents the DNA sequence of the small 16S-23S spacer region from Listeria seeligeri serovar 4A nr. 4268 (SEQ ID NO 118)
- Fig 44 : represents the partial DNA sequence of the large 16S-23S spacer region from partial sequence of the long spacer region of Listeria ivanovii CIP 7842 (SEQ ID NO 119)
- 10 Fig 45 : represents the DNA sequence of the large 16S-23S spacer region from Listeria monocytogenes IHE serovar 4B (SEQ ID NO 120)
- Fig 46 : represents the DNA sequence of the large 16S-23S spacer region from Listeria seeligeri serovar 4A nr. 4268 (SEQ ID NO 121)
- 15 Fig 47 : represents the DNA sequence of the 16S-23S spacer region from Chlamydia psittaci 6BC (SEQ ID NO 122)
- 20 Fig 48 : represents the DNA sequence of the 16S-23S spacer region from Chlamydia trachomatis (SEQ ID NO 123)
- Fig 49 : represents the DNA sequence of the 16S-23S spacer region from Mycoplasma genitalium (U. Gobel) (SEQ ID NO 124)
- 25 Fig 50 : represents the DNA sequence of the 16S-23S spacer region from Mycoplasma pneumoniae ATCC 29432 (SEQ ID NO 125)
- Fig 51 : represents the DNA sequence of the 16S-23S spacer region from Acinetobacter baumannii LMG 1041 (SEQ ID NO 126)
- 30 Fig 52 : represents the DNA sequence of the 16S-23S spacer region from Acinetobacter calcoaceticus LMG 1046 (SEQ ID NO 127)
- Fig 53 : represents the DNA sequence of the 16S-23S spacer region from Acinetobacter haemolyticus LMG 996 (SEQ ID NO 128)
- 35 Fig 54 : represents the DNA sequence of the 16S-23S spacer region from Acinetobacter johnsonii LMG 999 (SEQ ID NO 129)
- 40 Fig 55 : represents the DNA sequence of the 16S-23S spacer region from Acinetobacter junii LMG 998 (SEQ ID NO 130)
- Fig 56 : represents the DNA sequence of the 16S-23S spacer region from Brucella melitensis NIDO Biovar 1 (SEQ ID NO 131)
- 45 Fig 57 : represents the DNA sequence of the 16S-23S spacer region from Brucella suis NIDO Biovar 1 (SEQ ID NO 132)
- Fig 58 : represents the DNA sequence of one of the 16S-23S spacer region from Salmonella dublin (SEQ ID NO 133)
- 50 Fig 59 : represents the DNA sequence of one of the 16S-23S spacer region from Salmonella dublin (SEQ ID NO 134)
- 55 Fig 60 : represents the DNA sequence of one of the 16S-23S spacer region from Salmonella enteritidis (SEQ ID NO 135)
- Fig 61 : represents the DNA sequence of one of the 16S-23S spacer region from Salmonella enteritidis (SEQ ID

NO 136)

- Fig 62 : represents the DNA sequence of one of the 16S-23S spacer region from Salmonella typhimurium (SEQ ID NO 137)
- Fig 63: represents the DNA sequence of one of the 16S-23S spacer region from Salmonella typhimurium (SEQ ID NO 138)
- Fig 64 : represents the DNA sequence of one of the 16S-23S spacer region from Staphylococcus aureus strain UZG 5728 (SEQ ID NO 139)
- Fig 65 : represents the DNA sequence of one of the 16S-23S spacer region from Staphylococcus aureus strain UZG 6289 (SEQ ID NO 140)
- Fig 66 : represents the DNA sequence of one of the 16S-23S spacer region from Staphylococcus aureus strain UZG 6289 (SEQ ID NO 141)
- Fig 67 : represents the DNA sequence of one of the 16S-23S spacer region from Staphylococcus aureus strain UZG 6289 (SEQ ID NO 142)
- Fig 68 : represents the DNA sequence of one of the 16S-23S spacer region from Staphylococcus aureus strain UZG 6289 (SEQ ID NO 143)
- Fig 69 : represents the DNA sequence of one of the 16S-23S spacer region from Staphylococcus epidermidis strain UZG CNS41 (SEQ ID NO 144)
- Fig 70 : represents the DNA sequence of the 16S-23S spacer region from Streptococcus mitis UZG 2465 (SEQ ID NO 145)
- Fig 71 : represents the DNA sequence of the 16S-23S spacer region from Streptococcus pyogenes UZG 3671 (SEQ ID NO 146)
- Fig 72: represents the DNA sequence of the 16S-23S spacer region from Streptococcus sanguis UZG 1042 (SEQ ID NO 147)
- Fig 73 : represents the DNA sequence of the 16S-23S spacer region from Streptococcus saprophyticus UZG CNS46 (SEQ ID NO 148)
- Fig 74 : represents the DNA sequence of the 16S-23S spacer region from Streptococcus species UZG 536 (84) (SEQ ID NO 149)
- Fig 75 : represents the DNA sequence of the 16S-23S spacer region from Streptococcus species UZG 4341 (SEQ ID NO 150)
- Fig 76 : represents the DNA sequence of the 16S-23S spacer region from Streptococcus species UZG 457 (44B) (SEQ ID NO 151)
- Fig 77 : represents the DNA sequence of the 16S-23S spacer region from Streptococcus species UZG 97A (SEQ ID NO 152)
- Fig 78: represents the DNA sequence of the 16S-23S spacer region from Streptococcus species UZG 483 (76) (SEQ ID NO 153)
- Fig 79 : represents the DNA sequence of the 16S-23S spacer region from Brucella abortus NIDO Tulya biovar 3 (SEQ ID NO 154)
- Fig 80 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium ulcerans ITG 1837 and Mycobacterium marinum ITG 7732 (SEQ ID NO 157)

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- Fig 81 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium genavense ITG 8777 (SEQ ID NO 158)
- 5 Fig 82 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium genavense ITG 92-742 (SEQ ID NO 159)
- Fig 83 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium genavense ITG 9500 (SEQ ID NO 160)
- 10 Fig 84 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium simiae-like ITG 7379 (SEQ ID NO 161)
- Fig 85 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium simiae-like ITG 9745 (SEQ ID NO 162)
- 15 Fig 86 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium xenopi ITG 4986 (SEQ ID NO 163)
- Fig 87 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium simiae A ITG 4485 (SEQ ID NO 164)
- 20 Fig 88 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium simiae B ITG 4484 (SEQ ID NO 165)
- Fig 89 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium fortuitum ITG 4304 (SEQ ID NO 166)
- 25 Fig 90 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium kansasii ITG 6328 (SEQ ID NO 167)
- 30 Fig 91 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium kansasii ITG 8698 (SEQ ID NO 168)
- Fig 92 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium kansasii ITG 8973 (SEQ ID NO 169)
- 35 Fig 93 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium celatum ITG 94-123 (SEQ ID NO 170)
- Fig 94 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium haemophilum ITG 776 (SEQ ID NO 171)
- 40 Fig 95 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium haemophilum ITG 778 (SEQ ID NO 172)
- 45 Fig 96 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium haemophilum ITG 3071 (SEQ ID NO 173)
- Fig 97 : represents the DNA sequence of the 16S-23S spacer region from Mycobacterium chelonae ITG 94-330 and ITG 94-379 (SEQ ID NO 174)
- 50 Fig 98 : represents the DNA sequence of a 16S-23S spacer region from Yersinia enterocolitica strain P95 (SEQ ID NO 195)
- 55 Fig 99 : represents the DNA sequence of a 16S-23S spacer region from Yersinia enterocolitica strain P95 (SEQ ID NO 196)
- Fig 100 : represents the DNA sequence of the 16S-23S spacer region from Chlamydia trachomatis strain SSDZ

94 M 1961 (SEQ ID NO 197)

Fig 101 : represents the DNA sequence of a 16S-23S spacer region from Listeria-like isolate MB 405 (SEQ ID NO 213)

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Fig 102 : represents the DNA sequence of a 16S-23S spacer region from Listeria-like isolate MB 405 (SEQ ID NO 214)

10

Fig 103 : represents the DNA sequence of a 16S-23S spacer region from Listeria-like isolate MB 405 (SEQ ID NO 215)

TABLE LEGENDS

15	Table 1a:	List of all new probes originating from the 16S-23S rRNA spacer region
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100%

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Table 1a

	<u>PROBE</u>	<u>SEQUENCE</u>	<u>SEQ ID NO</u>
5	MYC-ICG-1	: ACTGGATAGTGGTTGCGAGCATCTA	1
	MYC-ICG-22	: CTTCTGAATAGTGGTTGCGAGCATCT	2
	MTB-ICG-1	: GGGTGCATGACAACAAAGTTGGCCA	3
10	MTB-ICG-2	: GACTTGTCCAGGTGTTGTCCCAC	4
	MTB-ICG-3	: CGGCTAGCGGTGGCGTGTCT	5
	MAI-ICG-1	: CAACAGCAAATGATTGCCAGACACAC	6
15	MIL-ICG-11	: GAGGGGTTCCCGTCTGTAGTG	7
	MIL-ICG-22	: TGAGGGGTTCTCGTCTGTAGTG	8
	MAC-ICG-1	: CACTCGGTGATCCGTGTGGA	9
20	MAV-ICG-1	: TCGGTCCGTCCGTGTGGAGTC	10
	MAV-ICG-22	: GTGGCCGGCGTTCATCGAAA	11
	MIN-ICG-1	: GCATAGTCCTAGGGCTGATGCGTT	12
25	MIN-ICG-2	: GCTGATGCGTTCGTCGAAATGTGTA	13
	MIN-ICG-22	: CTGATGCGTTCGTCGAAATGTGT	14
	MIN-ICG-222	: TGATGCGTTCGTCGAAATGTGT	15
30	MIN-ICG-2222	: GGCTGATGCGTTCGTCGAAATGTGTA	16
	MAL-ICG-1	: ACTAGATGAACCGCGTAGTCCTTGT	17
	MHEF-ICG-1	: TGGACGAAAACC GGTCACAA	18
35	MAH-ICG-1	: GTGTAATTCTTTTAACCTTGTGTGAAGTAAGTG	19
	MCO-ICG-11	: TGGCCGGCGTGTTCATCGAAA	20
	MTH-ICG-11	: GCACTTCAATTGGTGAAGT GCGAGCC	21
40	MTH-ICG-2	: GCGTGGTCTTCATGGCCGG	22
	MEF-ICG-11	: ACGCGTGGTCCTCGTGG	23
	MSC-ICG-1	: TCGGCTCGTCTGAGTGGTGT	24
45	MKA-ICG-1	: GATGCGTTGCTACGGGTAGCGT	25
	MKA-ICG-2	: GATGCGTTGCCTACGGGTAGCGT	26
	MKA-ICG-3	: ATGCGTTGCCCTACGGGTAGCGT	27
50	MKA-ICG-4	: CGGGCTCTGTCGAGAGTTGTC	28
	MCH-ICG-1	: GGTGTGGACTTGACTTCTGAATAG	29
	MCH-ICG-2	: CGGCAAAACGTCGGACTGTCA	30

	MCH-ICG-3	:	GGTGTGGTCCTGACTTATGGATAG	210
	MGO-ICG-1	:	AACACCCTCGGGTGCTGTCC	31
5	MGO-ICG-2	:	GTATGCGTTGTCGTCGCGGC	32
	MGO-ICG-5	:	CGTGAGGGGTACCGTCTGTAG	33
	MUL-ICG-1	:	GGTTTCGGGATGTTGTCACC	175
10	MGV-ICG-1	:	CGACTGAGGTCGACGTGGTGT	176
	MGV-ICG-2	:	GGTGTGAGCATTGAATAGTGGTTGC	177
	MGV-ICG-3	:	TCGGGCCGCGTGTCTCGTAAA	211
15	MXE-ICG-1	:	GTTGGGCAGCAGGCAGTAACC	178
	MSI-ICG-1	:	CCGGCAACGGTTACGTGTT	179
	MFO-ICG-1	:	TCGTTGGATGGCCTCGCACCT	180
20	MFO-ICG-2	:	ACTTGGCGTGGGATGCGGGAA	181
	MKA-ICG-5	:	CCCTCAGGGATTTCTGGGTGTTG	182
	MKA-ICG-6	:	GGACTCGTCCAAGAGTGTGTCC	183
25	MKA-ICG-7	:	TCGGGCTTGGCCAGAGCTGTT	184
	MKA-ICG-8	:	GGGTGCGAACAGCAAGCGA	185
	MKA-ICG-9	:	GATGCGTTGCCCTACGGG	186
30	MKA-ICG-10	:	CCCTACGGGTAGCGTGTCTTTG	187
	MML-ICG-1	:	CGGATCGATTGAGTGCTTGTCCC	188
	MML-ICG-2	:	TCTAAATGAACGCACTGCCGATGG	189
35	MCE-ICG-1	:	TGAGGGAGCCCGTGCCTGTA	190
	MHP-ICG-1	:	CATGTTGGCTTGATCGGTG	191
	PA-ICG 1	:	TGGTGTGCTCGTGATCCGAT	34
40	PA-ICG 2	:	TGAATGTTCGTGGATGAACATTGATT	35
	PA-ICG 3	:	CACTGGTGATCATTCAAGTCAAG	36
	PA-ICG 4	:	TGAATGTTCGT(G/A)(G/A)ATGAACATTGATTCTGGTC	37
45	PA-ICG 5	:	CTCTTCACTGGTGATCATTCAAGTCAAG	38
	LIS-ICG 1	:	CAAGTAACCGAGAATCATCTGAAAGTGAATC	39
	LMO-ICG 1	:	AAACAACCTTACTCGTAGAAGTAAATTGGTTAAG	40
50	LMO-ICG 2	:	TGAGAGGTTAGTACTCTCAGTATGTTGTT	41
	LMO-ICG 3	:	AGGCACTATGCTGAAGCATCGC	42
	LIV-ICG 1	:	GTTAGCATAAAATAGTAACATTATGACACAGAAC	43
55	LSE-ICG 1	:	AGTTAGCATAAGTAGTGTAACTATTTATGACACAGAAC	44

LISP-ICG 1	:	CGTTTCATAAGCGATCGCACGTT	212
5 CHTR-ICG 1	:	GGAAGAAGCCTGAGAAGGTTCTGAC	45
CHTR-ICG 2	:	GCATTATATGTAAGAGCAAGCATTCTATTCA	46
10 CHTR-ICG 3	:	GAGTAGCGTGGTGAGGACGAGA	47
CHPS-ICG 1	:	GGATAACTGTCTTAGGACGGTTGAC	48
15 MPN-ICG 1	:	ATCGGTGGTAAATTAAACCAAATCCCTGT	49
MPN-ICG 2	:	CAGTTCTGAAAGAACATTCCGCTTCTTC	50
15 MGE-ICG 1	:	CACCCATTAAITTTTCGGTGTAAAACCC	51
Mycoplasma-ICG	:	CAAAACTGAAAACGACAATCTTCTAGTTCC	52
20 STAU-ICG 1	:	TACCAAGCAAAACCGAGTGAATAAAGAGTT	53
STAU-ICG 2	:	CAGAAGATGCGGAATAACGTGAC	54
STAU-ICG 3	:	AACGAAGCCGTATGTGAGCATTGAC	55
STAU-ICG 4	:	GAACGTAACTTCATGTTAACGTTGACTTAT	56
25 ACI-ICG 1	:	GCTTAAGTGCACAGTGCTCTAAACTGA	57
ACI-ICG 2	:	CACGGTAATTAGTGTGATCTGACGAAG	58
BRU-ICG 1	:	CGTGCCGCCTCGTTCTCTTT	59
30 BRU-ICG 2	:	TTCGCTCGGGGTGGATCTGTG	60
BRU-ICG 3	:	GCGTAGTAGCGTTGCGTCGG	193
BRU-ICG 4	:	CGCAAGAAGCTTGCTCAAGCC	194
35 SALM-ICG 1	:	CAAAACTGACTTACGAGTCACGTTGAG	61
SALM-ICG 2	:	GATGTATGCTCGTTATTCCACGCC	62
40 STY-ICG 1	:	GGTCAAACCTCCAGGGACGCC	63
SED-ICG 1	:	GCGGTAAATGTGTGAAAGCGTTGCC	64
YEC-ICG 1	:	GGAAAAGGTACTGCACGTGACTG	198
45 YEC-ICG 2	:	GACAGCTGAAACTTATCCCTCCG	199
YEC-ICG 3	:	GCTACCTGTTGATGTAATGAGTCAC	200
CHTR-ICG 4	:	GAGTAGCGCGGTGAGGACGAGA	201

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Table 1b

<u>PRIMERS</u>	<u>SEQUENCE</u>	<u>SEQ ID NO</u>	
5			
MYC-P1	: TCCCTTGTGGCCTGTGTG	65	
MYC-P2	: TCCTTCATCGGCTCTCGA	66	
10	MYC-P3	: GATGCCAAGGCATCCACC	67
MYC-P4	: CCTCCCACGTCTTCATCG	68	
15	MYC-P5	: CCTGGGTTGACATGCACAG	192
CHTR-P1	: AAGGTTCTGACTAGGTTGGC	69	
20	CHTR-P2	: GGTGAAGTGCTTGCATGGATCT	70
LIS-P1	: ACCTGTGAGTTTCGTTCTCTC	71	
25	LIS-P2	: CTATTGTTCAAGTTGAGAGGTT	72
LIS-P3	: ATTTCCGTATCAGCGATGATAC	73	
LIS-P4	: ACGAACTAAAGGTTGTTTCT	74	
30	LIS-P5	: GAGAGGTTACTCTCTTATGTCAG	75
LIS-P6	: CTTTATGTCAGATAAAGTATGCAA	202	
LIS-P7	: CGTAAAAGGGTATGATTATTG	203	
35			
BRU-P1	: TCGAGAATTGGAAAGAGGTC	204	
BRU-P2	: AAGAGGTCGGATTATCCG	205	
40	BRU-P3	: TTGACTGCAAATGCTCG	206
BRU-P4	: TCTTAAAGCCGCATTATGC	207	
45			
YEC-P1	: CCTAATGATATTGATTGCG	208	
YEC-P2	: ATGACAGGTTAACCTTACCCC	209	
50			
EXAMPLE 1: <u>Pseudomonas aeruginosa</u>			

[0226] Pseudomonas aeruginosa is a significant human pathogen, usually in the context of serious underlying disease. It is also a major cause of nosocomial infections, which are characteristically prone to resistance to antimicrobial agents. This gram-negative, non-fermentative rod can be responsible for different clinical manifestations, like wound infections, bacteremia, respiratory and urinary tract infections, and is also a major cause of morbidity and mortality in patients with cystic fibrosis.

[0227] Pseudomonas species are currently differentiated based on growth characteristics and several biochemical features implying a time schedule of 24h to 72h to get a correct identification of the pathogen.

[0228] Already the development of monoclonal or polyclonal antibodies significantly improved the identification of Pseudomonas species. Recently however it has been shown that it is possible to detect organisms directly in clinical samples on a very sensitive and specific way using DNA probes with or without a prior amplification of the target DNA.

5 [0229] DNA probes to study Pseudomonas aeruginosa are already described and are mainly used for epidemiological typing (Ogle et al., 1987; Samadpour et al., 1988; McIntosh et al., 1992). However, none of these probes have been derived from the 16S-23S spacer.

10 [0230] The 16S-23S rRNA gene spacer region and a part of the 23S rRNA gene was amplified with conserved primers (upper primer: TGGGGTGAAGTCGTAACAAAGGTA , SEQ ID NO 155; lower primer: CCTTTCCCTCACGGTACTGGT, SEQ ID NO 156) using the polymerase chain reaction for the following species :

- 15
- Pseudomonas aeruginosa 5669
 - Pseudomonas alcaligenes LMG 1224^T
 - Pseudomonas fluorescens LMG 5167
 - Pseudomonas putida LMG 2232
 - Pseudomonas stutzeri LMG 2333^T
 - Pseudomonas pseudoalcaligenes LMG 1225^T

20 [0231] To facilitate cloning of the obtained amplicons a *NotI* recognition site was added to the lower primer. After purification and digestion of the fragment with *NotI*, the amplicon was cloned in a *EcoRV/NotI* digested pBluescript SK⁺ plasmid vector.

25 [0232] Sequencing of the 16S-23S rRNA gene spacer region was performed according the dideoxy-chain terminating chemistry either using double stranded plasmid DNA combined with primers located in the plasmid vector or directly on the PCR products after purification combined with internal PCR primers.

[0233] Fig. 36 to 40 represent the nucleotide sequence of the 16S-23S rRNA gene spacer regions from the different Pseudomonas species described above. For P. fluorescens only partial sequence information was obtained.

[0234] From the nucleic acid sequence of the spacer from P. aeruginosa strain 5669 five oligonucleotide-probes were chosen and chemically synthetized. The sequences of the oligonucleotides are the following :

30 PA1 = PA-ICG 1 : TGGTGTGCTGCGTGATCCGATA

PA2 = PA-ICG 2 : TGAATGTTCTGTGGATAACATTGATT

PA3 = PA-ICG 3 : CACTGGTGATCATTCAAGTCAAG

35 [0235] Specificity and sensitivity testing of the oligonucleotide-probes was carried out using a reverse hybridization assay. Genomic DNA of the different bacteria tested was amplified using biotinylated primers (idem primers as for cloning procedure, see above). The obtained amplicon, spanning the 16S-23S rRNA gene spacer region, was denatured and hybridized to a membrane-strip onto which the different oligonucleotide probes were immobilized in a line-wise fashion (LiPA). Hybridization was carried out in a mixture of 3xSSC (1xSSC = 0.15 M NaCl, 0.015 M sodium citrate, pH 7.0) and 20% formamide (FA) at a temperature of 50° C for one hour. Washing was done in the same mixture at the same temperature for 15 min.

40 [0236] Hybrids were detected using a streptavidine conjugate coupled to alkaline phosphatase and the probes were visualized through a precipitation reaction using NBT (nitroblue-tetrazolium) and BCIP (bromo-chloro-indolylphosphate).

45 [0237] The hybridization results obtained with probes PA1, PA2 and PA3 are given in table 4 and show that probes PA1 and PA3 were 100% specific for Pseudomonas aeruginosa and hybridized to all the strains tested. The hybridization signal with probe PA3 at 50° C was not optimal, so the oligonucleotide-probe was improved by adding some additional nucleotides to the specific probe. This newly designed probe is PA5.

50 PA5 = PA-ICG 5 : CTCTTCACTGGTGATCATTCAAGTCAAG

[0238] Hybridization experiments with probe PA5 proved that this probe also shows a 100% specificity and 100% sensitivity for P. aeruginosa.

55 [0239] Oligonucleotide-probe PA2 hybridized only to 5 out of 17 P. aeruginosa strains tested. Direct sequencing of the 16S-23S rRNA gene spacer region of the strains which did not hybridize to these probes, showed some heterogeneity between different strains. Two mismatches were seen in comparison to the first developed PA2 probe. To

overcome this heterogeneity between different strains in the region of probe PA2 a new probe PA4 was designed. This probe is degenerated at the position of the mismatches and some additional nucleotides were added to improve the hybridization signal at 50° C.

5

PA4 = PA-ICG 4 : TGAATGTTCGT(G/A)(G/A)ATGAACATTGATTCTGGTC

[0240] A 100% specificity and 100% sensitivity was obtained with this degenerated probe as is shown by the hybridization results. i

10

Table 2 :

Hybridization results for Pseudomonas						
	taxa tested	PA1	PA2	PA3	PA4	PA5
15	<u>Pseudomonas aeruginosa</u>	17/17	5/17	17/17	17/17	17/17
	<u>Pseudomonas alcaligenes</u>	0/1	0/1	0/1	0/1	0/1
	<u>Pseudomonas fluorescens</u>	0/1	0/1	0/1	0/1	0/1
20	<u>Pseudomonas putida</u>	0/1	0/1	0/1	0/1	0/1
	<u>Pseudomonas pseudoalcaligenes</u>	0/1	0/1	0/1	0/1	0/1
	<u>Pseudomonas stutzeri</u>	0/1	0/1	0/1	0/1	0/1
	<u>Pseudomonas cepacia</u>	0/1	0/1	0/1	ND	ND
25	<u>Neisseria gonorrhoeae</u>	0/1	0/1	0/1	ND	ND
	<u>Escherichia coli</u>	0/1	0/1	0/1	ND	ND
	<u>Bordetella pertussis</u>	0/1	0/1	0/1	ND	ND
	<u>Bordetella parapertussis</u>	0/1	0/1	0/1	ND	ND
	<u>Bordetella bronchiseptica</u>	0/1	0/1	0/1	ND	ND
30	<u>Mycobacterium tuberculosis</u>	0/1	0/1	0/1	ND	ND
	<u>Mycobacterium avium</u>	0/1	0/1	0/1	ND	ND
	<u>Moraxella catarrhalis</u>	0/4	0/4	0/4	ND	ND
	<u>Haemophilus influenzae</u>	0/2	0/2	0/2	ND	ND
	<u>Streptococcus pneumoniae</u>	0/3	0/3	0/3	ND	ND
35	<u>Acinetobacter calcoaceticus</u>	0/1	0/1	0/1	ND	ND
	<u>Staphylococcus aureus</u>	0/2	0/2	0/2	ND	ND
	(n/m: number of strains positive/number of strains tested)					
	(ND: not done)					

EXAMPLE 2: Mycobacterium

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[0241] A variety of mycobacterial species may be involved in serious human infectious disease. Notorious examples are *Mycobacterium tuberculosis* and *Mycobacterium leprae*. Recently other species such as *M. avium*, *M. intracellulare* and *M. kansasii* have been more frequently encountered as human pathogens especially in immunocompromised hosts.

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[0242] Consequently, laboratory diagnosis of mycobacterial infections should not be restricted to the *M. tuberculosis* complex but should ideally include most other clinically relevant mycobacterial species.

[0243] The identification and differentiation of pathogenic mycobacteria at the species level by conventional laboratory techniques is, in general, difficult and time-consuming.

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[0244] To overcome these problems DNA-techniques were implemented. The techniques described extended from straightforward DNA-probing to automated sequence analysis. Several approaches have been recently reported (Jonas et al., 1993; Frothingham and Wilson, 1993; Tomioka et al., 1993; Saito et al., 1989; Vaneechoutte et al., 1993; Telenti et al., 1993; Boddinghaus et al., 1990).

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[0245] However, these methods all have their particular disadvantages, and most of them still rely on culture. Moreover, and most importantly, none of these techniques allows for a simultaneous detection of the different clinically relevant mycobacterial species in a single test run. Besides, the differentiation of particular groups within the *Mycobacterium avium-intracellulare* complex is problematic and often even impossible.

[0246] To overcome the above-mentioned disadvantages, a LiPA-test was developed which allows for the simulta-

neous and reliable detection and differentiation of a number of *Mycobacterium* species and groups. The sets of probes used to achieve these goals were all derived from the 16S-23S rRNA spacer region. The methods used are analogous to those mentioned in example 1.

[0247] The 16S-23S rRNA spacer region, and part of the 16S and 23S rRNA flanking genes, was amplified by PCR with primers conserved for the genus *Mycobacterium*. At least one of the following primers located in the 16S gene were used as upper primers:

MYC-P1:	TCCCTTGTGGCCTGTGTG	(SEQ ID NO 65)
MYC-P5:	CCTGGGTTGACATGCACAG	(SEQ ID NO 192)

At least one of the following primers, located in the 23S gene, were used as lower primers for the amplification:

MYC-P2:	TCCTTCATCGGCTCTCGA	(SEQ ID NO 66)
MYC-P3:	GATGCCAAGGCATCCACC	(SEQ ID NO 67)
MYC-P4:	CCTCCCACGTCTTCATCG	(SEQ ID NO 68)

All the above mentioned primers amplified the spacer region of all *Mycobacterium* strains tested, except primer MYC-P2 which was not functional for *M. chelonae*. In order to enhance the sensitivity of the detection, a nested PCR was sometimes carried out, using P5 and P4 as outer primers and P1 and P3 as inner primers.

[0248] In order to be able to design and select the probes and probe combinations which fit our purpose, the 16S-23S rRNA spacer region of a number of mycobacterial strains was sequenced. The obtained sequences were compared to each other and to those already known from literature (e.g. Frothingham et al., 1993, 1994; Kampsell et al., 1992; Suzuki et al., 1988; EP-A-0395292; Van der Giessen et al., 1994;) or from publicly accessible data banks. The corresponding sequences are represented in fig.1 to 35 (SEQ ID NO 76 to SEQ ID NO 110).

[0249] The probes derived from these data were all adjusted in such a way that the desired hybridization-behaviour was obtained using unified hybridization and wash conditions (i.e. 3xSSC, 20% deionized formamide, 50°C). The set of adjusted probes used for hybridization to different mycobacterial strains is represented in table 1a, SEQ ID NO 1-33. Please note that the probe nomenclature used in this example is an abbreviated version of the one used in table 1a: i.e. the letters "ICG" have always been omitted. According to the specific hybridization pattern obtained, the strains tested could be assigned to one of the following species or species groups: *M. tuberculosis* complex, *M. avium*, *M. intracellulare* or *M. intracellulare* complex, *M. kansasii*, *M. chelonae* and *M. gordonaie*. The strains tested which belong to each group are summarized in Table 4. All strains were obtained from the Institute of Tropical Medicine, Antwerp, Belgium. The different probe-patterns obtained for each group are illustrated in Table 3, and are discussed in more detail hereafter.

40 *M. tuberculosis* complex

[0250] The *M. tuberculosis* complex harbours all strains belonging to *M. tuberculosis*, *M. bovis*, *M. africanum* and *M. microti*. The probes **Mtb1**, **Mtb2** and **Mtb3** hybridize with DNA originating from all *M. tuberculosis* complex strains tested. None of the other strains tested hybridized with these probes at the conditions used.

[0251] In addition, *M. tuberculosis* complex strains, as is the case with all other mycobacterial strains tested, hybridize with either the **myc1** or the **myc22** probe or both. The latter two probes are designed as general *Mycobacterium* probes, either alone or in combination with each other.

50 *M. avium/M. paratuberculosis*

[0252] All *M. avium* and *M. paratuberculosis* strains studied reveal an identical hybridization pattern with the set of probes. For this type of organisms positive hybridization signals are obtained with the probes **myc1/myc22**, **mai1**, **mai11**, **mav1**, **ma11** and **mav22**. The latter two probes hybridize exclusively with *M. avium* and *M. paratuberculosis* strains, and can thus be used as species-specific probes. Since the 16S-23S spacer sequences of *M. avium* isolates and *M. paratuberculosis* isolates are identical or nearly identical these two taxa cannot be discriminated from each other. This finding supports 16S rRNA sequencing data which indicate that *M. avium* and *M. paratuberculosis* should in fact be considered as belonging to one geno-species (Rogal et al., 1990), *M. avium* ssp. *avium* and *M. avium* ssp. *paratuber-*

culosis.

M. intracellulare and *M. intracellulare* complex (MIC)

- 5 [0253] MIC strains are genotypically highly related organisms, which, according to sequence data of the 16S-23S rRNA spacer region, belong to a distinct cluster which is separate from other *Mycobacterium* species. *M. avium* and *M. scrofulaceum* are their closest relatives. Almost all strains tested which are generally referred to as *M. avium* complex (MAC) strains (the former MAIS-complex) can be found in the MIC group. Thus, the MIC group defined in the current invention encompasses the MAC-type strains described by Frothingham and Wilson (1993) with the exception of MAC-10 G which appears to be *M. scrofulaceum*. Also *M. intracellulare* strains *sensu stricto* (*M. intracellulare* s.s.) are part of this cluster.
- [0254] Because this MIC group contains a quite large group of strains with, among them, subgroups showing different hybridization characteristics to the set of probes, a further subdivision into MIC-types was envisaged.
- 15 [0255] Type MIC 1 harbours *M. intracellulare* s.s., together with some other MAC-strains. All MIC 1 type isolates, without exception, hybridize to the following probes: **myc1/myc22, mali1 and mac1**. The following probes can be used to make further subdivisions within the MIC 1 group : **mil11, min1, min2 to 2222, mil22 and mhef1**.
- [0256] *M. intracellulare* *sensu stricto* strains (type MIC 1.1.a) can be distinguished from other subtypes in this group by virtue of probe **min1** which is positive only for this group of strains. All strains of type MIC 1.1.a strains are positive when tested with the *M. intracellulare* probe of the Gen-Probe Rapid Diagnostic system for MAC.
- 20 [0257] Type MIC 1.1.b and MIC 1.2 harbour strains which are highly related to *M. intracellulare*. They can be differentiated by using probes **mil11** and **mil22** (see Table 3). Further subdivision within these groups was not attempted although this could be achieved by using the probes : **min2, min22, min222** and **min2222**. Further subdivision might be of value for epidemiological reasons.
- [0258] Only two of our collection of strains tested group as MIC 2 strains. One of these strains is a "*Mycobacterium lufu*" strain (ITG 4755). The specific probe pattern generated by these strains is characterized by a positive hybridization signal with the following probes : **myc1/myc22, mali1, mil22, mah1 and mal1**. Variable hybridization results are obtained with probes **min2222, mac1** and **mhef1**. The other probes are negative. It is not unlikely that MIC 2 would eventually prove to be a heterogeneous group when more strains of this type are being identified. The variable probes may help in a further differentiation, if this would become relevant.
- 25 [0259] Type MIC 3 groups a fairly high number of MAC-strains which are rather remotely related to *M. intracellulare* s.s. strains and most other MAC-strains. This cluster should be regarded as distinct from *M. avium* and *M. intracellulare* on genotypical grounds. All
- [0260] MIC 3 subtypes hybridize to probes **myc1/myc22, mai1, mil22** and **mco1**. A positive signal with the latter probe (mco1) is characteristic for MIC 3 strains. Variable hybridization results are obtained with the following probes : **mac1 mhef1 and mah1**.
- 30 [0261] MIC 3 can be further subdivided into four subtypes by using three probes : **mh11, mth2** and **mef11**. Probe **mth2** is specific for type MIC 3.1 which encompasses a group of highly related MAC-strains isolated from immunocompromised human beings.
- [0262] Most MIC 3 strains are located in the MIC 3.1 subtype. Eventually species status may be assigned to this group of strains, as might also be the case for other groups of MAC strains, yet unnamed. In subtypes MIC 3.4, MIC 3.3 and MIC 3.2 only two, one and one strain are found respectively in our collection of strains tested.
- 35 [0263] Type MIC 4 is a collection of "MAIS" strains (including *M. malmoense*) which are remotely related to *M. intracellulare*. The only probe of the above-described set which hybridizes to MIC 4, apart from the general myc1/myc22 probes, is the mai1 probe. This probe shows a broad specificity, hybridizing also with *M. avium*, *M. intracellulare* and other MIC strains and *M. scrofulaceum*.

M. scrofulaceum

- 40 [0264] All *M. scrofulaceum* strains tested reveal an identical hybridization pattern with the set of probes. A positive signal with probe **msc1** is unique to *M. scrofulaceum* strains. The only other probes with a positive signal for this species are evidently **myc1/myc22** and also **mai1**.

M. kansasii

- 45 [0265] Probes **mka3** and **mka4** are specific for *M. kansasii*; i.e. a distinct positive signal is obtained on the LiPA strip when amplified DNA from the *M. kansasii* strains is used in the hybridization whilst with all other organisms tested the signal is absent. Although the sequences of probes **mka1** and **mka2** are not absolutely complementary to the target sequence (3 and 1 mismatches, respectively), these probes also proved to be useful since they hybridized exclusively

to *M. kansasii* DNA and not to any other mycobacterial DNA tested under the conditions used (50°C, 3xSSC, 20% formamide). This illustrates that probes not necessarily have to match perfectly to the target to be useful, and that modifications in sequence and length may be allowed up to a certain degree.

5 ***M. chelonae***

[0266] The species *M. chelonae* encompasses *M. chelonae* ssp. *chelonae* and *M. chelonae* ssp. *abscessus* strains. The spacer region was sequenced for one strain of each subspecies and small differences were noticed (SEQ ID NO 103 and SEQ ID NO 102). Probes **mch1** and **mch2** hybridize to both strains. All other probes are negative for these 2 strains except for **myc1/myc22**.

[0267] Upon testing of probes **mch1** and **mch2** with 2 additional *M. chelonae* strains not mentioned in table 4, i.e. *M. chelonae* 94-379 and *M. chelonae* 94-330, both obtained from the Institute of Tropical Medecine in Antwerp, Belgium, it appeared that they did not hybridize to probe **mch1**. This was confirmed by sequencing the spacer region of these two strains (SEQ ID NO 184). Cluster analysis of the spacer region with other mycobacteria revealed that *M. chelonae* strains can be subdivided in two groups. A third probe **mch3** was designed to specifically detect this second group of strains, to which 94-379 and 94-330 belong.

[0268] This illustrates that the use of DNA probes derived from the 16S-23S rRNA spacer region can be helpful in differentiating different groups of strains, which belong to the same species according to the classical identification methods, and possibly can be used to detect and describe new species within the mycobacteria. In this case **mch2** detects all *M. chelonae* strains, whereas **mch1** and **mch3** differentiate between different subgroups.

M. gordonaie

[0269] The five *M. gordonaie* strains tested all hybridize to probe **mgo5**. Positive hybridization signals are also obtained with probes **myc1/myc22**, and some *M. gordonaie* strains also hybridize to probes **mgo1** and **mgo2**.

other mycobacterial species

[0270] Strains belonging to other mycobacterial species than those mentioned above only hybridize to the general probes **myc1/myc22**. This indicates that these strains most probably belong to the genus *Mycobacterium*, but do not belong to one of the species or groups which can be specifically identified by using one or more of the other probes described.

[0271] In conclusion we can state that, according to the particular combinations of probes of the invention used, DNA probe tests at different levels can be provided.

[0272] When all probes are used in one and the same LiPA-test, differentiation at the species level as well as sub-typing of certain groups of mycobacteria can be achieved. However, the probe-assembly on one strip could be restricted to those probes which are species-specific; in that case identification is performed at the species level. A further reduction of the number of probes on the strip might lead to the specific detection of only one or just a few species. Obviously, LiPA strips can be designed which solely attempt to subtype strains, e.g. those belonging to the *M. intracellulare* complex (MIC). Depending on the particular needs of the laboratoria performing diagnosis and/or typing of mycobacteria, all these different applications might be of value. However, it is clear that by using a combination of probes in a LiPA-format the amount of information obtained as to the identity of the organisms present in the clinical sample, is considerably increased as compared to DNA probe tests using only a single probe. For some groups, or at least for further subdivision of some groups, a single probe uniquely hybridizing to this (sub)group could not be designed. In that case only probe-patterns are able to provide the information needed. For these applications the LiPA is an advantageous format.

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Table 3 :
Different probe patterns obtained for mycobacterial (sub)species

Mycobacterium	mycl myc22	mtb1 mtb2 mtb3	mai1	mtb11	mav1 mav22	nim1	min222	min2	min2222	min22	mac1
<i>M. tuberculosis</i>	+	+	-	-	-	-	-	-	-	-	-
<i>M. bovis</i>		-	+	+	-	-	-	-	-	-	-
<i>M. avium</i>			-	-	-	-	-	-	-	-	-
<i>M. paratuberculosis</i>			-	-	-	-	-	-	-	-	-
<i>MIC 1.1.a</i>			+	+	-	-	-	-	-	-	-
<i>MIC 1.1.b</i>			+	+	-	-	-	-	-	-	-
<i>MIC 1.2</i>			+	+	-	-	-	-	-	-	-
<i>MIC 2</i>			+	-	-	-	-	-	-	-	-
<i>MIC 3.4</i>			+	+	+	-	-	-	-	-	-
<i>MIC 3.3</i>			+	+	+	-	-	-	-	-	-
<i>MIC 3.1</i>			+	+	+	-	-	-	-	-	-
<i>MIC 3.2</i>			+	+	+	-	-	-	-	-	-
<i>MIC 4</i>			+	-	-	-	-	-	-	-	-
<i>M. scrofulaceum</i>			+	-	-	-	-	-	-	-	-
<i>M. kansasii</i>			-	-	-	-	-	-	-	-	-
<i>M. chelonae</i>			-	-	-	-	-	-	-	-	-
<i>M. gordoniæ</i>			-	-	-	-	-	-	-	-	-
<i>Mycobacterium sp.</i>			-	-	-	-	-	-	-	-	-

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Table 3: continued

Mycobacterium	mco1	mth11	mth2	meff1	meff1	maff1	maff1	mse1	mka1,2,3,4	mch1,2,3	mgo1,2	mgo5
<i>M. tuberculosis</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>M. bovis</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>M. avium</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>M. paratuberculosis</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>MIC 1.1.a</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>MIC 1.1.b</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>MIC 1.2</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>MIC 2</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>MIC 3.4</i>	+	+	+	+	+	+	+	-	-	-	-	-
<i>MIC 3.3</i>	+	+	+	+	+	+	+	-	-	-	-	-
<i>MIC 3.1</i>	+	+	+	+	+	+	+	-	-	-	-	-
<i>MIC 3.2</i>	+	+	+	+	+	+	+	-	-	-	-	-
<i>MIC 4</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>M. scrofulaceum</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>M. kansasii</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>M. chelonae</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>M. gordonaiae</i>	-	-	-	-	-	-	-	-	-	-	-	-
<i>Mycobacterium sp.</i>	-	-	-	-	-	-	-	-	-	-	-	-

w : weak / v : very weak / + : + or - variable according to the strain tested

Table 4

Mycobacteria strains tested in LIPA		
	species/group	strain numbers from Institute of Tropical Medecine Antwerp (except those between parentheses)
5	M. tuberculosis complex	7602, 8004, 8017, 8647, 8872, 9081, 9129, 9173, 9517, (ATCC 27294), 8324, 8428
10	M.avium/ M. paratuberculosis	1101, 1983, 2070, 2074, 4176, 4189, 4191, 4193, 4197, 4204, 4386, 4991, 5872, 5874, 5884, 5887, 5893, 5894, 5897, 5903, 5904, 5905, 5927, 5983, 8180, 8750, (ATCC 25291), <u>M. paratub.</u> (316F), (2E)
15	M. intracellulare (MIC 1.1.a)	4199, 4208, 5701, 5880, 5906, 5908, 5909, 5913, 5915, 5917, 5918, 5920, 5921, 5924, 5925, 5929, 8713, 8717, 8718, 8720, 8721, 8722, 8732, 8740, 8741, 8742, 8744, 8747, 8749
20	MIC 1.1.b	8694, 8745, 8754 8708 5513, 8743 8054, 8190
25	MIC 1.2	8710, 8711, 8712, 8714, 8715, 8716, 8725, 8729, 8733, 8737, 8746, 8751, 8752 5919 8695 8748
30	MIC 2	5922 4755 (M. lufu)
35	MIC 3.4	1815 8707
40	MIC 3.3	5620
45	MIC 3.1	925, 926, 1329, 1788, 1794, 1812, 1818, 2069, 2073, 2076, 4541, 4543, 5074, 5280, 5789, 7395, 8739, 8753 8738
50	MIC 3.2	5765
	M. scrofulaceum	4979, 4988, 5907, 8706, 8726, 8727, 8735, (MB022), (MB023), (MB024)
	M. kansasii	4987, (ATCC 22478)
	M. chelonae	4975, 9855
	M. gordoneae	7703, 7704, 7836, 7838, 8059
	MIC 4	8723, 8724 8757 4842 (M. malmoense)
	other mycobacterial species	7732 (M. marinum), 94-123 (M. celatum), 778 (M. haemophilum), 8777 (M. genavense), 4484 (M. siniae), 4986 (M. xenopi), 4304 (M. fortuitum), 1837 (M. ulcerans)

EXAMPLE 3: Listeria

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[0273] Listeria species are a group of Gram-positive rods widely spread in nature. Within this group it seems that only L. monocytogenes is pathogenic to humans and animals. L. monocytogenes is the causative agent of listeriosis, giving rise to meningitis, abortions, encephalitis and septicemia. Immunocompromised individuals, newborn infants

and pregnant women are high risk groups for this foodborn disease. Most cases have been caused by the consumption of food of animal origin, particularly soft cheeses. Therefore, the presence of *L. monocytogenes* should be excluded from food. For safety measurements, in some countries, the absence of all *Listeria* species is required in food products.

[0274] The classical identification method for *L. monocytogenes* in dairy products involves an enrichment culture for 48 h and subsequently colony forming on selective agar medium for 48 h followed by a whole set of biochemical and morphological assays (Farber and Peterkin, 1991). This procedure could be very much simplified by the use of gene probes.

[0275] Several DNA probes are already described for the identification of *L. monocytogenes*. Some probes are derived from genes responsible for the pathogenicity of the organism, for instance the listeriolysin O gene (Datta et al., 1993) or the invasion-associated-protein (iap) (Bubert et al., 1992).

[0276] A commercially available identification system, based on a specific 16S rRNA probe, was introduced by Gen-Probe (Herman and De Ridder, 1993; Ninet et al., 1992).

[0277] These specific probes are used as confirmation assays on colonies obtained after enrichment and plating on selective agar medium.

[0278] Recently several publications reported on the use of the polymerase chain reaction to amplify the target region for the DNA probes, which can shorten the time of the assay without interfering with the specificity and the sensitivity of the assay. Different primer sets are described that can specifically amplify *L. monocytogenes* DNA. These primer sets were derived from the listeriolysin O gene (Golstein Thomas et al., 1991), and the iap gene (Jaton et al., 1992).

[0279] We used the 16S-23S rRNA gene spacer region as the target for the development of a genus-specific probe for *Listeria* and a probe specific for *Listeria monocytogenes*.

[0280] Using conserved primers derived from the 3' end of the 16S rRNA and the 5' end of the 23S rRNA (sequences are given in example 1) the spacer region was amplified using the polymerase chain reaction and subsequently cloned in a suitable plasmid vector following the same procedures as in example 3.

[0281] Two amplicons differing in length (800 bp and 1100 bp) were obtained. Both PCR fragments were cloned for the following *Listeria* species:

- *Listeria monocytogenes*, serovar 4b, IHE (Instituut voor Hygiëne en Epidemiologie, Belgium)
- *Listeria ivanovii* CIP 78.42 (Collection Nationale de Cultures de Microorganisms de l'Institut Pasteur, France)
- *Listeria seeligeri* serovar 4a, nr. 42.68 (Bacteriologisches Institut, Südd, Versuchs- und Forschungsanstalt für Milchwirtschaft Weihenstephan, Germany)

[0282] The sequence of the spacer region between the 16S and 23S rRNA gene was determined using the cloned material originating from the 800 bp PCR fragment and this was done for the three described *Listeria* species. Fig. 41 to 43 show the sequences of the different short spacer regions obtained. The sequence of this short spacer region of *L. monocytogenes* was also retrieved from the EMBL databank (LMRGSPCR).

[0283] Based on this sequence information, following oligonucleotides for species-specific detection were chosen and chemically synthesized :

40 LMO-ICG-1 : AAACAACCTTACTTCGTAGAAGTAAATTGGTTAAG

LMO-ICG-2 : TGAGAGGTAGTACTTCTCAGTATGTTGTC

LSE-ICG-1 : AGTTAGCATAAGTAGTGTAACTATTTATGACACAAG

45 LIV-ICG-1 : GTTAGCATAAATAGTAACTATTTATGACACAAGTAAC

Also, a genus specific probe for *Listeria* was designed:

50 LIS-ICG-1 : CAAGTAACCGAGAATCATCTGAAAGTGAATC

The oligonucleotide-probes were immobilized on a membrane strip and following reverse hybridization with biotinylated PCR fragments, the hybrids were visualized using a precipitation reaction. The hybridization results of different *Listeria* species are summarized in table 5.

Table 5

Species	n	LIS1	LMO1	LMO2	LSE1 1	LIV1
<u>L. monocytogenes</u>	1	+	+	+	-	-
<u>L. seeligeri</u>	2	+	+	±	-	±
<u>L. ivanovii</u>	3	+	±	-	±	+
<u>L. welshimeri</u>	3	+	+	±	-	-
<u>L. innocua</u>	2	+	+	+	-	-

[0284] These hybridization results show that probe LIS1 can detect all described Listeria species, but also that the species-specific probes cross-hybridize to each other. Hence, from this short spacer region probes with sufficient specificity could not be found.

[0285] For Listeria monocytogenes the 16S-23S rRNA gene spacer was also determined originating from the 1100 bp fragment. Fig. 45 shows the sequence obtained for this species. This sequence information was also obtained for L. seeligeri (see fig. 46) and partial sequence information of the large spacer region was obtained for L. ivanovii (see fig. 44).

[0286] Based on sequence alignment with L. seeligeri following oligonucleotide-probe was chosen to specifically detect L. monocytogenes.

LMO-ICG-3 : AGGCACATATGCTTGAAGCATCGC

[0287] Initial hybridization results (not shown) indicated that no cross-hybridization with other Listeria species was seen with this L. monocytogenes probe LMO3, and that all Listeria strains used hybridized to the general probe LIS1.

[0288] The oligonucleotide-probes, LIS1 for detection of all Listeria species and LMO3 for specific detection of L. monocytogenes, were immobilized on a membrane strip and hybridized to labeled amplicons, containing the 16S-23S rRNA spacer region, derived from different organisms. The hybridization results are shown in the following table.

[0289] An excellent specificity and sensitivity were obtained for probes LMO3 and LIST respectively at the species and genus level.

Table 6

Taxa tested	n	LIS1	LMO3
<u>Listeria monocytogenes</u>	44	+	+
<u>Listeria ivanovii</u>	10	+	-
<u>Listeria seeligeri</u>	11	+	-
<u>Listeria welshimeri</u>	16	+	-
<u>Listeria innocua</u>	23	+	-
<u>Listeria murrayi</u>	3	+	-
<u>Listeria grayi</u>	2	+	-
<u>Brochotrix thermosphacta</u>	1	-	-
<u>Brochotrix campestris</u>	1	-	-
<u>Bacillus cereus</u>	3	-	-
<u>Bacillus brevis</u>	2	-	-
<u>Bacillus coagulans</u>	1	-	-
<u>Bacillus pumilis</u>	1	-	-
<u>Bacillus macerans</u>	1	-	-
<u>Bacillus lentinus</u>	1	-	-
<u>Bacillus firmus</u>	2	-	-
<u>Bacillus subtilis</u>	2	-	-
<u>Bacillus megantum</u>	1	-	-
<u>Enterococcus faecalis</u>	1	-	-
<u>Enterococcus faecium</u>	1	-	-

Table 6 (continued)

Taxa tested	n	LIS1	LMO3
<u>Enterococcus durans</u>	1	-	-
<u>Lactococcus lactis</u>	3	-	-
<u>Lactococcus casei</u>	1	-	-
<u>Escherichia coli</u>	1	-	-
<u>Hafnia halvei</u>	1	-	-
<u>Agrobacterium tumefaciens</u>	2	-	-
<u>Mycoplasma dimorpha</u>	1	-	-
<u>Clostridium tyrobutyricum</u>	1	-	-
<u>Clostridium perfringens</u>	1	-	-
<u>Clostridium sporogenes</u>	1	-	-
<u>Clostridium acetobutylicum</u>	1	-	-
<u>Brucella abortus</u>	1	-	-
<u>Brucella suis</u>	1	-	-
<u>Brucella melitensis</u>	1	-	-
<u>Staphylococcus aureus</u>	1	-	-
<u>Salmonella typhimurium</u>	1	-	-
<u>Salmonella enteritidis</u>	1	-	-
<u>Yersinia enterocolitica</u>	1	-	-

n: number of strains tested

[0290] These two probes can be used for the detection of Listeria species and Listeria monocytogenes directly on food samples or after enrichment of the samples in liquid broth. In both cases amplification problems can occur with the conserved primerset due to the enormous background flora in these samples.

[0291] To circumvent this problem, we designed several sets of primers derived from the 16S-23S rRNA spacer regions of Listeria species.

[0292] Primers LIS-P1 and LIS-P2 are upper primers, whereas LIS-P3 and LIS-P4 are lower primers. These primersets amplify the smaller 16S-23S rRNA spacer region as well as the larger spacer of Listeria species (except L. grayi and L. murrayi). If needed these primers can be used in a nested PCR assay where LIS-P1/LIS-P4 are the outer primers and LIS-P2/LIS-P3 are the inner primers.

[0293] For the specific detection of Listeria monocytogenes probe LMO-ICG-3 was designed and derived from the large 16S-23S rRNA spacer region. In order to specifically amplify only this large spacer region for an improved detection of this pathogen directly in samples a set of primers was derived from the part of sequence information from the large 16S-23S rRNA spacer region that is not present in the smaller rRNA spacer. For this aim, primers LIS-P5 and LIS-P6 are used as the upper primers and LIS-P7 is used as the lower primer.

LIS-P1	:	ACCTGTGAGTTTCGTTCTTCTC	71
LIS-P2	:	CTATTTGTTCAGTTTGAGAGGTT	72
LIS-P3	:	ATTTTCCGTATCAGCGATGATAC	73
LIS-P4	:	ACGAAGTAAAGGTTGTTTCT	74
LIS-P5	:	GAGAGGTTACTCTTTATGTCAG	75
LIS-P6	:	CTTTTATGTCAGATAAAGTATGCAA	202
LIS-P7	:	CGTAAAAGGGTATGATTATTTG	203

[0294] During the evaluation of the probes for Listeria spp. an organism was isolated from cheese that resembled Listeria according to the classical determination methods. This isolate (MB 405) showed the following characteristics (similar to Listeria spp.): Gram positive, growth on Oxford and Tryptic Soy Agar, catalase positive. The only difference with the Listeria spp. was the motility, which was negative.

[0295] Using the conserved primers as described in example 1 in order to amplify the 16S-23S rRNA spacer region of this isolate MB 405, the same amplicon pattern was obtained with this strain as with Listeria spp. Hybridization of the amplicon showed that there was no signal obtained with any of the probes for Listeria spp.

[0296] Sequencing of the 16S rRNA of isolate MB 405 and subsequent comparison with Listeria spp. and relatives showed that the organism was more closely related to Listeria spp. than to any other species described in the literature until now. Taxonomical studies will show if this isolate does or does not belong to the genus Listeria. This isolate, and subsequently isolated organisms from the same type, are referred to in this application as Listeria like organisms.

[0297] Isolate MB 405 seemed to contain at least 3 different 16S-23S rRNA spacer regions which were cloned and sequenced. Following alignment with Listeria spp. an oligonucleotide-probe was chosen to specifically detect Listeria-like strains:

LISP-ICG-1 : CGTTTCATAAGCGATCGCACGTT

15 Reverse hybridization reactions of this probe with the 16S-23S rRNA spacer regions of Listeria spp. showed that there was no cross-hybridization.

EXAMPLE 4: Chlamydia trachomatis

20 [0298] Chlamydia trachomatis is a small obligate intracellular gram-negative bacterium, which has 15 serovars (A-K, Ba, L1, L2, and L3) distinguished by the major outer membrane protein (MOMP) and contains a cryptic plasmid required for intracellular growth. The A-K and Ba serovars constitute the trachoma biovar, while the L1, L2, and L3 serovars constitute the LGV biovar.

25 [0299] Serovars A, B, Ba, and C are commonly associated with trachoma, the leading cause of preventable blindness worldwide. The D-K serovars are found mainly in sexually transmitted infections and are the major cause of cervicitis and pelvic inflammatory disease in women, and urethritis and epididymitis in men. Serovars L1, L2 and L3 are involved in lymphogranuloma venereum, a rare sexually transmitted disease.

30 [0300] Cell culture is regarded as the benchmark method for laboratory diagnosis, although specimen viability is difficult to maintain during transport and laboratory techniques are time-consuming and technically demanding. Therefore, a number of more rapid test kits were developed, such as an enzyme-linked immunosorbent assay, and direct fluorescent-antibody staining. However, none of these immunoassays have been shown to have high levels of sensitivity or specificity.

35 [0301] A nonisotopic DNA probe assay (Gen-Probe PACE; Woods et al., 1990) that detects chlamydial rRNA is commercially available. Recently, the polymerase chain reaction (PCR) method has been used for detection of Chlamydia infections. Detection was targeted at either the cryptic plasmid (Loeffelholz et al., 1992), or the *omp1* gene, which encodes for the major outer membrane protein (Taylor-Robinson et al., 1992). Compared with other techniques, PCR has higher sensitivity and specificity (Ossewaarde et al., 1992). None of these assays make use of DNA probes derived from the 16S-23S rRNA gene spacer region.

40 [0302] For a Chlamydia trachomatis L2 and a Chlamydia psittaci 6BC strain, a part of the ribosomal RNA cistron, containing the 16S-23S rRNA spacer region was amplified using conserved primers (see example 1) and subsequently cloned in a plasmid vector. The 16S-23S rRNA spacer region was sequenced using the dideoxynucleotide terminating chemistry.

[0303] The sequence of the spacer region of both Chlamydia species is shown in fig. 47 to 48.

45 [0304] Based on this sequence information, following oligonucleotide-probes were chemically synthesized :

CHTR-ICG-1 : GGAAGAACGCTGAGAAGGTTCTGAC

CHTR-ICG-2 : GCATTATATGTAAGAGCAAGCATTCTATTCA

50 CHTR-ICG-3 : GAGTAGCGTGGTGAGGACGAGA

CHPS-ICG-1 : GGATAACTGTCTAGGACGGTTGAC

55 [0305] The oligonucleotide-probes were immobilized in a line-wise fashion on a membrane strip and subsequently used in a reverse hybridization assay with biotinylated PCR products, containing the 16S-23S rRNA spacer region, as target.

[0306] Hybridizations were done in a solution of 3xSSC and 20% formamide (FA) at a temperature of 50°C.

[0307] The hybridization results with the different probes are shown in the following table.

Table 7

Strains tested	CHTR1	CHTR2	CHTR3	CHPS1
<u>Chlamydia trachomatis</u> L2	+	+	+	-
<u>Chlamydia psittaci</u> 6BC	-	-	-	+
<u>Chlamydia psittaci</u> CP	-	-	-	+
<u>Chlamydia psittaci</u> TT	-	-	-	+
<u>Haemophilus ducreyi</u> CIP 542	-	-	-	-
<u>Haemophilus influenzae</u> NCTC 8143	-	-	-	-
<u>Neisseria gonorrhoeae</u> NCTC 8375	-	-	-	-
<u>Moraxella catarrhalis</u> LMG 5128	-	-	-	-
<u>Escherichia coli</u> B	-	-	-	-
<u>Streptococcus pneumoniae</u> S92-2102	-	-	-	-

[0308] As shown in the table at a hybridization temperature of 50°C the probes CHTR1, CHTR2 and CHTR3 are specific for Chlamydia trachomatis and probe CHPS1 is specific for Chlamydia psittaci.

[0309] Several clinical isolates, obtained from the SSDZ, Delft, Netherlands, identified as Chlamydia trachomatis using conventional methods were tested in a reverse hybridization assay with the different oligonucleotide-probes. All

Chlamydia trachomatis specific probes gave a positive hybridization signal and none of the isolates reacted with the Chlamydia psittaci probe. For some clinical isolates the CHTR2 probe reacted significantly weaker than CHTR1 or CHTR3. The spacer region of one of these isolates (94 M 1961) was sequenced (SEQ ID NO 197) and the sequence

revealed one mismatch with the spacer sequence of strain L2. An additional probe (CHTR4) was derived from this new spacer sequence:

CHTR-ICG-4 : GAGTAGCGCGGTGAGGGACGAGA (SEQ ID NO 201)

This probe gives a stronger hybridization signal than CHTR2 with some clinical isolates from Chlamydia trachomatis. It can be used alone, or in combination with the CHTR2 probe (e.g. both probes applied in one LiPA-line).

[0310] In order to develop very sensitive assays for the detection of Chlamydia trachomatis directly in clinical specimens a specific primerset was derived from the 16S-23S rRNA spacer region, CHTR-P1 (upper primer) and CHTR-P2 (lower primer), amplifying specifically the spacer region of Chlamydia species.

CHTR-P1 : AAGGTTTCTGACTAGGTTGGGC 69

CHTR-P2 : GGTGAAGTGCTTCATGGATCT 70

EXAMPLE 6: Mycoplasma pneumoniae and Mycoplasma genitalium

[0311] Mycoplasmas are a group of the smallest prokaryotes known that are able to grow in cell-free media, lack a cell wall, and have very small genomes with a low G+C content. More than 100 different species have been isolated from humans, animals, plants, and insects.

[0312] In humans, mycoplasmas have been recognized either as pathogenic organisms or as commensals. The best known pathogen is Mycoplasma pneumoniae, the causative agent of primary atypical pneumonia, especially in children and young adults. The diagnosis of M. pneumoniae has been based on the direct isolation by the culture method or on the detection of specific antibodies against M. pneumoniae in the patient's serum.

[0313] Another pathogen, first isolated from urethral specimens from patients with nongonococcal urethritis, has been described as Mycoplasma genitalium. This mycoplasma has several properties in common with M. pneumoniae. Both species are pathogenic, and both possess the capability to adhere to erythrocytes, various tissue cells, glass, and plastic surfaces. Furthermore, M. genitalium and M. pneumoniae share antigens, giving rise to extensive cross-reactions in serological tests. The observation that M. genitalium could also be found in respiratory tract specimens from patients with pneumonia and isolated from a mixture with M. pneumoniae has raised questions to the possible pathogenicity of M. genitalium.

[0314] Since cultivation of both species is time-consuming and serology lacks specificity, more rapid and more specific assays were developed to identify these mycoplasmas. The use of hybridization assays with DNA probes was described for these species, but despite good specificities these tests do not allow the detection of low levels of M. pneumoniae or M. genitalium. So more recently, DNA hybridization techniques were developed using the polymerase chain reaction. M. pneumoniae-specific PCR assays have been reported using the P1 adhesin gene (Buck et al., 1992) and the 16S rRNA gene (Kuppeveld et al., 1992). Specific PCR assays for M. genitalium were described using sequences from the adhesin gene and the 16S rRNA gene.

[0315] The spacer sequences of clinical isolates of M. pneumoniae and M. genitalium (obtained from U. Gobel, University of Freiburg, Germany) were determined. They are shown in fig. 49 to 50. The sequences show some differences to those from other strains of the same species deposited in the EMBL databank (MPMAC and MGG37 respectively). Based on this information four probes were derived: one general Mycoplasma probe, two M. pneumoniae specific, and one M. genitalium specific probe:

15 Mycoplasma-ICG: CAAAACGTAAACGACAATCTTCTAGTTCC
 MPN-ICG-1: ATCGGTGGTAAATTAAACCCAAATCCCTGT
 MPN-ICG-2: CAGTTCTGAAAGAACATTCCGCTTCTTC
 20 MGE-ICG-1: CACCCATTAATTTTCGGTGTAAAACCC

[0316] The probes were applied to LiPA strips and hybridized under standard conditions (3X SSC, 20% formamide at 50°C) to amplified spacer material from four M. pneumoniae strains, one M. genitalium strain and twenty-two non-Mycoplasma species strains. The general probe hybridized only to the five Mycoplasma strains tested, while the specific probes hybridized only to strains of the species for which they were designed.

EXAMPLE 7: Other mycobacterial species

[0317] With the steady improvement of laboratory techniques the information on the systematics and clinical significance of the so called "potentially pathogenic environmental mycobacteria" increased rapidly. With the emergence of newly recognized diseases, additional syndromes associated with different mycobacterial species have emerged and have assumed major importance.

[0318] In order to extend the LiPA test for the simultaneous detection of different mycobacterial species as described in example 2, a new set of DNA probes was designed to specifically identify the following species : Mycobacterium ulcerans, Mycobacterium genavense, Mycobacterium xenopi, Mycobacterium simiae, Mycobacterium fortuitum, Mycobacterium malmoense, Mycobacterium celatum and Mycobacterium haemophilum.

[0319] These probes were derived from the 16S-23S rRNA spacer region sequence. For the above mentioned species this information was obtained through direct sequencing of PCR products or after cloning of the PCR-amplified spacer region. The sequences obtained are represented in fig. 80 to 97, and in fig. 38 for M. malmoense.

[0320] The sequences of the spacer region of the above-mentioned mycobacterial species were compared and aligned to those already described in example 2 or in publicly available sources. From the regions of divergence, species-specific DNA probes were designed. The probes were selected and designed in such a way that the desired hybridization behaviour (i.e. species-specific hybridization) was obtained under the same conditions as those specified for the other mycobacterial probes mentioned in example 2, i.e. 3X SSC, 20% deionized formamide, 50°C. This allows simultaneous detection of at least two, and possibly all, of the mycobacterial species described in the current invention.

[0321] The following oligonucleotide probes were designed from the spacer region sequence of respectively M. ulcerans, M. genavense, M. xenopi, M. simiae, M. fortuitum, M. malmoense, M. celatum and M. haemophilum:

MUL-ICG-1 : GGTTTCGGGATGTTGTCCCACC
 5 MGV-ICG-1 : CGACTGAGGTCGACGTGGTGT
 MGV-ICG-2 : GGTGTTGAGCATTGAATAGTGGTTGC
 MXE-ICG-1 : GTTGGGCAGCAGGCAGTAACC
 10 MSI-ICG-1 : GCCGGCAACGGTTACGTGTT
 MFO-ICG-1 : TCGTTGGATGGCCTCGCACCT

15 MFO-ICG-2 : ACTTGGCGTGGGATGCGGGAA
 MML-ICG-1: CGGATCGATTGAGTGCTTGTCCC
 MML-ICG-2: TCTAAATGAACGCAGTGCCTGATGG
 20 MCE-ICG-1: TGAGGGAGCCCGTGCCTGTA
 MHP-ICG-1: CATGTTGGGCTTGATCGGGTGC

25 [0322] The probes were immobilized on a LiPA strip and hybridized with amplified biotinylated material derived from a set of representative mycobacterial species as described in example 2. Amplification of the spacer region was carried out by PCR using a primer set as described in example 2. The different strains used for specificity testing are shown in table 8 together with the hybridization results obtained. The strains were obtained from the collection of the Institute for Tropical Medicine, Antwerp, Belgium.

30 [0323] The probes tested (MSI-ICG1, MXE-ICG-1, MFO-ICG-1, MFO-ICG-2, MML-ICG-1, MML-ICG-2, MCE-ICG-1 and MHP-ICG-1) specifically detected *M. simiae*, *M. xenopi*, *M. fortuitum*, *M. malmoense*, *M. celatum* and *M. haemophilum* respectively and showed no cross-hybridization with the other mycobacterial species tested. Thus, these probes allow a specific detection of mycobacterial species which were not further identifiable using the set of DNA probes described in example 2. *M. malmoense* was classified in example 2 as a "MIC 4"-type, while the other species mentioned above were only hybridizing to the general probes MYC1/MYC22 for the genus *Mycobacterium*, and were thus classified in example 2 as "other mycobacterial species".

35 [0324] All tested *M. genavense* isolates reacted with MGV-ICG1 and MGV-ICG2, and not with MSI-ICG1 designed for *M. simiae*, closely related to *M. genavense*. A group of "intermediate" organisms, situated in between *M. simiae* and *M. genavense*, were received from the Tropical Institute of Medecine, Antwerp, where they were classified as "*M. simiae* - like" (strains 4358, 4824, 4833, 4844, 4849, 4857, 4859, 7375, 7379, 7730, 9745, 94-1228). These strains reacted only with probe MGV-ICG2 and not with probe MSI-ICG1 which specifically detects *M. simiae* strains *sensu stricto*. Sequencing of the 16S-23S rRNA spacer region of two of these "*M. simiae*-like" isolates (strains 7379 and 9745) (see SEQ ID NO 161 and 162) confirmed that they were more closely related to *M. genavense* than to *M. simiae*. A new probe MGV-ICG3 was designed to specifically detect this group of organisms, which possibly belong to a new species.

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MGV-ICG 3 : TCAGGGCCCGTGTTCGTCAA

50 [0325] This illustrates again that the use of DNA probes derived from the 16S-23S spacer region can be helpful in differentiating different groups of strains, which are also found indeterminate by classical taxonomic criteria. The use of these DNA probes may possibly lead to the description of new (sub)species within mycobacteria. In this case, the MGV-1 probe would react only with *M. genavense* strains *sensu stricto*, MGV-3 probe would react only with the intermediate "*M. simiae*-like" strains, and MGV-2 probe would detect both types of strains.

55 [0326] The probe MUL-ICG-1 reacted with all *M. ulcerans* strains tested, but also showed cross-hybridization with *M. marinum* strain ITG 7732. Sequencing of the spacer region of this *M. marinum* strain indeed revealed an identical sequence to that of *M. ulcerans* strain 1837 (see fig. 80). Further differentiation between *M. marinum* and *M. ulcerans* can be done using a probe from the 16S-rRNA gene of *M. ulcerans*, part of which is co-amplified with the spacer region when primers MYC P1 -P5 are used for amplification. A species-specific 16S rRNA probe for *M. ulcerans*, which can

work under the same hybridization conditions as the spacer probes for mycobacterium species differentiation, is for example :

5 TGGCCGGTGCAAAGGGCTG

(SEQ ID NO 216)

[0327] The above paragraph shows that, although it is preferable to use probes derived from the spacer region, it is also possible, and sometimes necessary, to combine the spacer probes with probes derived from other gene sequences, e.g. the 16S rRNA gene. Here again, these additional probes are selected such that they show the desired hybridization characteristics under the same hybridization and wash conditions as the spacer probes.

[0328] For M. kansasii, additional strains to the ones mentioned in example 2 have been tested with probes MKA-ICG-1, 2, 3 and 4 described in example 2. Since none of these probes was entirely satisfactory, additional probes were designed for M. kansasii detection. Therefor, the spacer region of some of the additional M. kansasii strains ITG 6328, 8698 and 8973 was sequenced (see fig.90 to 92). These strains were also obtained from the Institute of Tropical Medicine in Antwerp, Belgium. Apparently, M. kansasii strains constitute a quite heterogeneous group, with remarkable differences in the spacer sequence between different strains. Additional probes MKA-ICG-5, 6, 7, 8, 9 and 10 were designed, all hybridizing again under the same conditions as those earlier described, i.e. 3X SSC, 20% deionized formamide, 50°C. The probes were tested with a collection of test strains obtained from the Institute of Tropical Medicine, Antwerp, Belgium, and results are shown in table 8.

[0329] None of the M. kansasii probes hybridizes with a species other than M. kansasii, as far as tested. However, due to the heterogeneous character of this species, none of the M. kansasii probes hybridizes with all M. kansasii strains. The different M. kansasii probes recognize different strains of M. kansasii. This differential hybridization may be of clinical significance. On the other hand, if detection of all M. kansasii strains is desirable, a combination of different M. kansasii probes can be envisaged.

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Table 8: additional mycobacterial probes

species/type	strain	MUL ICG-1	MGV ICG-1 2 3	MXE ICG-1 ICG-2	MFO ICG-1 ICG-2	MSI ICG-1	MML ICG-1 ICG-2	MCB ICG-1	MHP ICG-1
<i>M. tuberculosis</i>	8004	-	-	-	-	-	-	-	-
<i>M. avium</i>	5887	-	-	-	-	-	-	-	-
<i>M. intracellulare</i>	5915	-	-	-	-	-	-	-	-
	5913	-	-	-	-	-	-	-	-
<i>MIC 3.1 strain</i>	1812	-	-	-	-	-	-	-	-
<i>MIC-4 strain</i>	8724	-	-	-	-	-	-	-	-
<i>M. scrofulaceum</i>	4979	-	-	-	-	-	-	-	-
<i>M. kansasii</i>	4987	-	-	-	-	-	-	-	-
	2795	-	-	-	-	-	-	-	-
	6228	-	-	-	-	-	-	-	-
	6362	-	-	-	-	-	-	-	-
	8698	-	-	-	-	-	-	-	-
	8973	-	-	-	-	-	-	-	-
	8974	-	-	-	-	-	-	-	-
	8971	-	-	-	-	-	-	-	-
<i>M. ulcerans</i>	1837	+	-	-	-	-	-	-	-
	3129	+	-	-	-	-	-	-	-
	5114	+	-	-	-	-	-	-	-
	5115	+	-	-	-	-	-	-	-
<i>M. marinum</i>	7732	+	-	-	-	-	-	-	-
<i>M. malmense</i>	4832	-	-	-	-	-	-	-	-
	4842	-	-	-	-	-	-	-	-
<i>M. gordonaie</i>	7703	-	-	-	-	-	-	-	-

Table 8 continued

\pm = negative reaction, $+$ = positive reaction, w = weak reaction, \pm = variable reaction, blanc = non tested

Table 8 continued

species/type	strain	MKA ICG-3	MKA ICG-4	MKA ICG-5	MKA ICG-6	MKA ICG-7	MKA ICG-8	MKA ICG-9	MKA- ICG-10
<i>M. tuberculosis</i>	8004	-	-	-	-	-	-	-	-
<i>M. avium</i>	5887	-	-	-	-	-	-	-	-
<i>M. intracellulare</i>	5915 5913	-	-	-	-	-	-	-	-
<i>MIC 3.1 strain</i>	1812	-	-	-	-	-	-	-	-
<i>MIC-4 strain</i>	8724	-	-	-	-	-	-	-	-
<i>M. scrofulaceum</i>	4979	-	-	-	-	-	-	-	-
<i>M. kansassii</i>	4987 2795 6238 6362 8698 8973 8974 8971	+ + + + - - - -	+ + + + - - - -	- - - - - - - -	- - - - +	- - - - - - - -	- - - - - - - -	- - - - - - - -	- - - - - - - -
<i>M. ulcerans</i>	1837 3129 5114 5115								
<i>M. marinum</i>	7732	-	-	-	-	-	-	-	-
<i>M. malmoense</i>	4832 4842	-	-	-	-	-	-	-	-
<i>M. gordonaie</i>	7703	-	-	-	-	-	-	-	-

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Table 8 continued

<i>M. chelonae</i>	4975 9855 94-330 94-379							
<i>M. celatum</i>	94-123							
<i>M. haemophilum</i>	778 3071							
<i>M. genavense</i> and <i>M. simiae-like</i>	8777 9745 92-742 7379 9500							
<i>M. simiae</i>	4484 4485							
<i>M. xenopi</i>	4986							
<i>M. fortuitum</i>	4304							

55 EXAMPLE 8: Brucella

- [0330] Brucellosis is a very widespread and economically important zoonosis which also affects humans.
 [0331] For the identification of Brucella spp., mainly bacteriological and immunological detection techniques are

being used. These tests are time-consuming and often give false-positive results. Quick and reliable identification methods are being developed, mainly based on DNA amplification and hybridization.

[0332] Specific detection of Brucella spp. based on the amplification of a 43 kDa outer membrane protein (Fekete A. et al., 1990) or of a part of the 16S rRNA gene (Herman and De Ridder, 1992) were already described.

5 [0333] In order to develop specific DNA probes and primers for the detection of Brucella spp. we analyzed the 16S-23S rRNA gene spacer region. Using conserved primers (sequences are given in example 1) the spacer region was amplified and subsequently cloned into the Bluescript SK+ vector following the same procedures as in example 1. The obtained amplicon of about 1400 bp in length was cloned for the following Brucella species:

- 10 - Brucella abortus NIDO Tulya biovar 3 (SEQ ID NO 154)
 - Brucella melitensis NIDO biovar 1 (SEQ ID NO 131)
 - Brucella suis NIDO biovar 1 (SEQ ID NO 132)

15 *Hind* III digestion of the constructs, followed by subcloning of the obtained fragments (n=3) facilitated the sequencing of the spacer region for the three described Brucella spp..

Fig. 56, 57 and 79 represent the sequences of the spacer regions obtained for the above-mentioned strains of respectively Brucella melitensis, Brucella suis and Brucella abortus.

Due to the high homology of these spacer region sequences between different Brucella species, no species-specific DNA probes were deduced from this sequence information, and only genus-specific probes were designed.

20 [0334] For this purpose, the following probes were chemically synthesized:

BRU-ICG 1 : CGTGCCGCCTCGTTCTCTTT
 BRU-ICG 2 : TTTCGCTTCGGGGTGGATCTGTG
 BRU-ICG 3 : GCGTAGTAGCGTTGCGTCGG
 BRU-ICG 4 : CGCAAGAAGCTTGCTCAAGCC

30 The oligonucleotides were immobilized on a membrane strip and following reverse hybridization with biotinylated PCR fragments, the hybrids were visualized using a precipitation reaction. The hybridization results of the immobilized probes with different Brucella spp. and related organisms are represented in the table 9.

35 [0335] These hybridization results show that probes BRU-ICG 2, BRU-ICG 3 and BRU-ICG 4 are specific for Brucella spp. and can be used in a reverse hybridization assay for detection of these pathogens. Probe BRU-ICG 1 cross-hybridizes with Ochrobactrum antropi and Rhizobium loti strains, which are two taxonomically highly related organisms, but which are not expected to be present in the same sample material as used for Brucella detection.

[0336] As described in previous examples (e.g. 3 and 4) also for Brucella specific primers were chosen from the 16S-23S rRNA spacer region, in order to specifically amplify the spacer region from Brucella strains.

40 [0337] BRU-P1 and BRU-P2 are used as upper primers, while BRU-P3 and BRU-P4 are used as lower primers. When used in a nested PCR assay the combination BRU-P1/BRU-4 is the outer primerset whereas the combination BRU-P2/BRU-P3 is the inner primerset.

45	BRU-P1 : TCGAGAATTGGAAAGAGAGTC	204
	BRU-P2 : AAGAGGTCGGATTATCCG	205
	BRU-P3 : TTTCGACTGCAAATGCTCG	206
50	BRU-P4 : TCTTAAAGCCGCATTATGCG	207

TABLE 9

TAXA TESTED	n	BRU-ICG 1	BRU-ICG 2	BRU-ICG 3	BRU-ICG 4
<u>Brucella abortus</u>	6	+	+	+	+
<u>Brucella suis</u>	3	+	+	+	+

TABLE 9 (continued)

	TAXA TESTED	n	BRU-ICG 1	BRU-ICG 2	BRU-ICG 3	BRU-ICG 4
5	<u>Brucella melitensis</u>	4	+	+	+	+
	<u>Brucella ovis</u>	2	+	+	+	+
	<u>Brucella cams</u>	2	+	+	+	+
	<u>Brucella neotomae</u>	1	+	+	+	+
	<u>Phyllobacterium rubiacearum</u>	1	-	-	NT	NT
10	<u>Ochrobactrum anthropi</u>	8	+	-	-	-
	<u>Agrobacterium tumefaciens</u>	2	-	-	NT	NT
	<u>Agrobacterium rhizogenes</u>	1	-	-	NT	NT
	<u>Mycoplana dimorpha</u>	1	-	-	NT	NT
	<u>Rhizobium loti</u>	1	+	-	-	-
15	<u>Rhizobium meliloti</u>	1	-	-	NT	NT
	<u>Rhizobium leguminosarum</u>	1	-	-	NT	NT
	<u>Bradyrhizobium japonicum</u>	1	-	-	NT	NT
	<u>Brochothrix thermosphacta</u>	1	-	-	NT	NT
	<u>Brochothrix campestris</u>	1	-	-	NT	NT
20	<u>Bacillus cereus</u>	3	-	-	NT	NT
	<u>Bacillus brevis</u>	2	-	-	NT	NT
	<u>Bacillus coagulans</u>	1	-	-	NT	NT
	<u>Bacillus pumilis</u>	1	-	-	NT	NT
	<u>Bacillus macerans</u>	1	-	-	NT	NT
25	<u>Bacillus lenth</u>	1	-	-	NT	NT
	<u>Bacillus firmus</u>	2	-	-	NT	NT
	<u>Bacillus subtilis</u>	2	-	-	NT	NT
	<u>Bacillus megantum</u>	1	-	-	NT	NT
	<u>Enterococcus faecalis</u>	1	-	-	NT	NT
30	<u>Enterococcus faecium</u>	1	-	-	NT	NT
	<u>Enterococcus durans</u>	1	-	-	NT	NT
	<u>Lactobacillus lactis</u>	3	-	-	NT	NT
	<u>Lactobacillus casei</u>	1	-	-	NT	NT
	<u>Leuconostoc lactis</u>	1	-	-	NT	NT
35	<u>Escherichia coli</u>	1	-	-	NT	NT
	<u>Hafnia halvei</u>	1	-	-	NT	NT
	<u>Clostridium tyrobutyricum</u>	1	-	-	NT	NT
	<u>Clostridium perfringens</u>	1	-	-	NT	NT
	<u>Clostridium sporogenes</u>	1	-	-	NT	NT
40	<u>Clostridium acetobutylicum</u>	1	-	-	NT	NT
	<u>Staphylococcus aureus</u>	1	-	-	NT	NT
	<u>Salmonella enteritidis</u>	1	-	-	NT	NT
	<u>Yersinia enterocolitica</u>	1	-	-	NT	NT
	<u>Listeria monocytogenes</u>	1	-	-	NT	NT
45	<u>Listeria ivanovii</u>	1	-	-	NT	NT
	<u>Listeria seeligeri</u>	1	-	-	NT	NT
	<u>Listeria welshimeri</u>	1	-	-	NT	NT
	<u>Listeria innocua</u>	1	-	-	NT	NT
	<u>Listeria murrayi</u>	1	-	-	NT	NT
50	<u>Listeria grayi</u>	1	-	-	NT	NT
	NT = Not tested	n = number of strains tested				

EXAMPLE 9: *Staphylococcus aureus*

[0338] *Staphylococcus aureus* is the staphylococcal species most commonly associated with human and animal infections. *Staphylococcus aureus* strains have been identified as important etiologic agents in both community-acquired and nosocomial infections. Recently nosocomial infection with methicillin-resistant *S. aureus* (MRSA) appear to be increasingly prevalent in many countries. The strains belonging to this species are also causative agents of food spoilage and poisoning.

[0339] In order to discriminate in a fast and specific way *S. aureus* strains from other staphylococci, the use of molecular techniques based on DNA probes and/or PCR were already described in the literature. Examples of target genes used for the development of these DNA based assays are the 16S rRNA gene (De Buyser et al., 1992; Geha et al, 1994), the *mecA* gene (Ubukata et al., 1992; Shimaoka et al., 1994) and the *nuc* gene (Brakstad et al., 1992; Chesneau et al., 1993).

[0340] As a target for the development of specific DNA probes we chose the 16S-23S rRNA gene spacer region. Amplification using conserved primers derived from the 16S and the 23S rRNA genes (sequences, see example 1) showed that the pattern obtained was not similar in all *S. aureus* strains tested. A lot of variation was seen in either the number of fragments obtained and in the size of these different fragments.

[0341] One spacer region from strain UZG 5728 and four spacer regions (differing in length) from strain UZG 6289 were cloned into Bluescript SK+ vector and subsequently sequenced. The sequences are represented in fig. 64 to fig. 68 (SEQ ID NO 139 to SEQ ID NO 143). For the development of specific DNA probes these different spacer regions were compared to each other and to the spacer region derived from *Staphylococcus epidermidis* strain UZG CNS41 (SEQ ID NO 144).

[0342] The following probes were chemically synthesized :

25 STAU-ICG 1 : TACCAAGCAAAACCGAGTGAATAAAGAGTT

STAU-ICG 2 : CAGAAGATGCCGAATAACGTGAC

STAU-ICG 3 : AACGAAGCCGTATGTGAGCATTGAC

30 STAU-ICG 4 : GAACGTAACCTCATGTTAACGTTGACTTAT

[0343] The oligonucleotides were immobilized on a membrane strip and following reverse hybridization with biotinylated PCR fragments, the hybrids were visualized using a colorimetric precipitation reaction.

[0344] The hybridization results of the immobilized probes with different *Staphylococcus* spp. and non-staphylococcal organisms are represented in Table 10.

[0345] These hybridization results show that only probes STAU-ICG 3 and STAU-ICG 4 are specific for *Staphylococcus aureus* strains. Probe STAU-ICG 1 reacts with all *Staphylococcus* spp. tested and probe STAU-ICG 2 cross-hybridizes with the *S. lugdinensis* strain.

Neither probe STAU-ICG 3 nor probe STAU-ICG 4 detects all *S. aureus* strains tested, but when both probes are used simultaneously in a LiPA assay, all *S. aureus* strains tested hybridize with one of these probes or with both.

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Table 10

Strains tested	n	STAU-ICG 1	STAU-ICG 2	STAU-ICG 3	STAU-ICG 4
<i>Staphylococcus aureus</i>	13	+			
<i>Staphylococcus aureus</i>	10	+			
<i>Staphylococcus aureus</i>	3	+			
<i>staphylococcus aureus</i>	1	+			
<i>Staphylococcus epidermidis</i>	11	+			
<i>Staphylococcus saprophyticus</i>	1	+			
<i>Staphylococcus haemolyticus</i>	1	+			
<i>Staphylococcus capitis</i>	1	+			
<i>Staphylococcus lugdunensis</i>	1	+			
<i>Staphylococcus hominis</i>	1	+			
<i>Bordetella pertussis</i>	1	+			
<i>Bordetella parapertussis</i>	1				
<i>Bordetella bronchiseptica</i>	1				
<i>Mycobacterium tuberculosis</i>	1				
<i>Mycobacterium avium</i>	1				
<i>Moraxella catarrhalis</i>	4				
<i>Haemophilus influenzae</i>	2				
<i>Streptococcus pneumoniae</i>	3				
<i>Pseudomonas cepacia</i>	1				
<i>Pseudomonas aeruginosa</i>	3				
<i>Acinetobacter calcoaceticus</i>	1				

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SEQUENCE LISTING

(1) GENERAL INFORMATION:

5

(i) APPLICANT:

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10

15 (ii) TITLE OF INVENTION: SIMULTANEOUS DETECTION, IDENTIFICATION AND DIFFERENTIATION OF EUBACTERIAL TAXA USING A HYBRIDIZATION ASSAY

(iii) NUMBER OF SEQUENCES: 216

20

(iv) COMPUTER READABLE FORM:

- (A) MEDIUM TYPE: Floppy disk
- (B) COMPUTER: IBM PC compatible
- (C) OPERATING SYSTEM: PC-DOS/MS-DOS
- (D) SOFTWARE: PatentIn Release #1.0, Version #1.25 (EPO)

25

(2) INFORMATION FOR SEQ ID NO: 1:

30

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 25 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

35

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

ACTGGATAGT GGTTGCGAGC ATCTA

25

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(2) INFORMATION FOR SEQ ID NO: 2:

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(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 26 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

55

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

55

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

CTTCTGAATA GTGGTTGCGA GCATCT

26

5 (2) INFORMATION FOR SEQ ID NO: 3:

- (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 25 base pairs
(B) TYPE: nucleic acid
10 (C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

15 (iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

GGGTGCATGA CAACAAAGTT GGCCA

25

(2) INFORMATION FOR SEQ ID NO: 4:

- 25 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 24 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

30 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

35 (iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

40 GACTTGTTC AGGTGTTGTC CCAC

24

(2) INFORMATION FOR SEQ ID NO: 5:

- 45 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 21 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

50 (iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

CGGCTAGCGG TGGCGTGTTC T

21

5 (2) INFORMATION FOR SEQ ID NO: 6:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 26 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

15 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:

25 CAACAGCAAA TGATTGCCAG ACACAC

26

28 (2) INFORMATION FOR SEQ ID NO: 7:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 21 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

30 (ii) MOLECULE TYPE: cDNA

35 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

38 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:

42 GAGGGGTTCC CGTCTGTAGT G

21

45 (2) INFORMATION FOR SEQ ID NO: 8:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 22 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

48 (ii) MOLECULE TYPE: cDNA

52 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

TGAGGGGTTCTCGTCTGTAG TG

22

(2) INFORMATION FOR SEQ ID NO: 9:

- 5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 21 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear
- 10 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iv) ANTI-SENSE: NO

15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:

20 CACTCGGTCTGATCCGTGTGG A

21

(2) INFORMATION FOR SEQ ID NO: 10:

- 25 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 21 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear
- 30 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iv) ANTI-SENSE: NO

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:

TCGGTCCGTC CGTGTGGAGT C

21

40 (2) INFORMATION FOR SEQ ID NO: 11:

- 45 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 20 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear
- 50 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iv) ANTI-SENSE: NO

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

GTGGCCGGCG TTCATCGAAA

20

(2) INFORMATION FOR SEQ ID NO: 12:

- 5 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 25 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

15

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

GCATAGTCCT TAGGGCTGAT GCGTT

25

20 (2) INFORMATION FOR SEQ ID NO: 13:

- 25 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 25 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

30 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

GCTGATGCGT TCGTCGAAAT GTGTA

25

(2) INFORMATION FOR SEQ ID NO: 14:

40

- (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 23 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

45

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

50

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:

55

CTGATGCGTT CGTCGAAATG TGT

23

(2) INFORMATION FOR SEQ ID NO: 15:

- 5 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 22 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

15

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:

TGATGCGTTC GTCGAAATGT GT

22

20 (2) INFORMATION FOR SEQ ID NO: 16:

- 25 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 27 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

30

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

GGCTGATGCG TTCTGTCGAAA TGTGTAA

27

(2) INFORMATION FOR SEQ ID NO: 17:

- 40 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 24 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

45

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

50

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

ACTAGATGAA CGCGTAGTCC TTGT

24

55

(2) INFORMATION FOR SEQ ID NO: 18:

(ii) MOLECULE TYPE: cDNA

10 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:

TGGACGAAAAA CCCGGGTGCAC AA

22

(2) INFORMATION FOR SEO ID NO: 19:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 38 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

30 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 19:

35 GTGTAATTTC TTTTTAACT CTTGTGTGA AGTAAGTG

38

(2) INFORMATION FOR SEQ ID NO: 20:

40 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 21 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

50

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

TGGCCGGCGT GTTCATCGAA A

21

55 (2) INFORMATION FOR SEQ ID NO: 21:

- 5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 26 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

10 (iii) HYPOTHETICAL: NO

10 (iii) ANTI-SENSE: NO

15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

GCACTTCAAT TGGTGAAGTG CGAGCC

26

20 (2) INFORMATION FOR SEQ ID NO: 22:

- 20 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 19 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

25 (ii) MOLECULE TYPE: cDNA

25 (iii) HYPOTHETICAL: NO

25 (iii) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:

GCGTGGTCTT CATGGCCGG

19

35 (2) INFORMATION FOR SEQ ID NO: 23:

- 35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 18 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA

40 (iii) HYPOTHETICAL: NO

40 (iii) ANTI-SENSE: NO

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

ACGGCGTGGTC CTTCGTGG

18

55 (2) INFORMATION FOR SEQ ID NO: 24:

- 55 (i) SEQUENCE CHARACTERISTICS:

- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

10 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:

15 ATGCGTTGCC CTACGGGTAG CGT

23

(2) INFORMATION FOR SEQ ID NO: 28:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 22 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

25 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

CGGGCTCTGT TCGAGAGTTG TC

22

(2) INFORMATION FOR SEQ ID NO: 29:

- 35
- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

45 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

50 GGTGTGGACT TTGACTTCTG AATAG

25

(2) INFORMATION FOR SEQ ID NO: 30:

- 55
- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 21 base pairs
 - (B) TYPE: nucleic acid

- (C) STRANDEDNESS: single
(D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

CGGCACAAACG TCGGACTGTCA A

21

15

(2) INFORMATION FOR SEQ ID NO: 31:

- (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 20 base pairs
(B) TYPE: nucleic acid
20 (C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

25

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:

AACACCCCTCG GGTGCTGTCC

20

(2) INFORMATION FOR SEQ ID NO: 32:

35

- (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 21 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

40

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

45

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:

50

GTATGCGTTG TCGTTCGCGG C

21

(2) INFORMATION FOR SEQ ID NO: 33:

55

- (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 22 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

5 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

CGTGAGGGGT CATCGTCTGT AG

22

15 (2) INFORMATION FOR SEQ ID NO: 34:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 21 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
20 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

25 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:

30 TGGTGTGCTG CGTGATCCGA T

21

(2) INFORMATION FOR SEQ ID NO: 35:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 26 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
35 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

40 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

45

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:

TGAATGTTCG TGGATGAACA TTGATT

26

50 (2) INFORMATION FOR SEQ ID NO: 36:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 23 base pairs
(B) TYPE: nucleic acid
55 (C) STRANDEDNESS: single
(D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

CACTGGTGAT CATTCAAGTC AAG

23

(2) INFORMATION FOR SEQ ID NO: 37:

- 15 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 33 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:

30 TGAATGTTCG TVVATGAACA TTGATTCTG GTC

33

(2) INFORMATION FOR SEQ ID NO: 38:

- 35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 29 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

40 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

CTCTTCACT GGTGATCATT CAAGTCAAG

29

(2) INFORMATION FOR SEQ ID NO: 39:

- 50 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 31 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

CAAGTAACCG AGAACATCT GAAAGTGAAT C

31

(2) INFORMATION FOR SEQ ID NO: 40:

15 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 36 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:

AAACAACCTT TACTTCGTAG AAGTAAATTG GTTAAG

36

30 (2) INFORMATION FOR SEQ ID NO: 41:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 32 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

40 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

45

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

TGAGAGGTTA GTACTTCTCA GTATGTTGT TC

32

(2) INFORMATION FOR SEQ ID NO: 42:

50

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 23 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

55

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:

10 AGGCACATATG CTTGAAGCAT CGC

23

(2) INFORMATION FOR SEQ ID NO: 43:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 38 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

20 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:

GTTCAGCATAA ATAGGTAACT ATTTATGACA CAAGTAAC

38

30 (2) INFORMATION FOR SEQ ID NO: 44:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 36 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

40 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:

45 AGTTAGCATA AGTAGTGTAA CTATTATGA CACAAG

36

(2) INFORMATION FOR SEQ ID NO: 45:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 26 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

55 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 45:

GGAAGAAGCC TGACAAGGTT TCTGAC

26

10

(2) INFORMATION FOR SEQ ID NO: 46:

15

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 33 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

20

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:

GCATTTATAT GTAAGAGCAA GCATTCTATT TCA

33

30

(2) INFORMATION FOR SEQ ID NO: 47:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 22 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

35

(ii) MOLECULE TYPE: cDNA

40

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

45

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:

GAGTAGCGTG GTGAGGACGA GA

22

50

(2) INFORMATION FOR SEQ ID NO: 48:

55

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 26 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (iii) HYPOTHETICAL: NO

5
 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:

GGATAACTGT CTTAGGACGG TTTGAC

26

10 (2) INFORMATION FOR SEQ ID NO: 49:

15 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 30 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

20

(iii) ANTI-SENSE: NO

25

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 49:

ATCGGTGGTA ATTAAACCC AAATCCCTGT

30

(2) INFORMATION FOR SEQ ID NO: 50:

30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 30 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

35

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 50:

CAGTTCTGAA AGAACATTTC CGCTTCTTTC

30

45

(2) INFORMATION FOR SEQ ID NO: 51:

50

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 30 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

55

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 51:

CACCCATTAA TTTTTTCGGT GTTAAAACCC

30

(2) INFORMATION FOR SEQ ID NO: 52:

10 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 31 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: cDNA
(iii) HYPOTHETICAL: NO

20 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 52:

25 CAAAACTGAA AACGACAATC TTTCTAGTTC C

31

(2) INFORMATION FOR SEQ ID NO: 53:

30 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 30 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA
(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:
TACCAAGCAA AACCGAGTGA ATAAAGAGTT

30

45 (2) INFORMATION FOR SEQ ID NO: 54:

50 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 23 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

55 (iii) ANTI-SENSE: NO

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 54:

CAGAAAGATGC GGAATAACGT GAC

23

(2) INFORMATION FOR SEQ ID NO: 55:

10 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 26 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

20

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 55:

AACCGAAGCCG TATGTGAGCA TTTGAC

26

(2) INFORMATION FOR SEQ ID NO: 56:

30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 31 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

35 (iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 56:

GAACGTAACT TCATGTTAAC GTTTGACTTA T

31

(2) INFORMATION FOR SEQ ID NO: 57:

45 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 27 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

55 (iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:

5 GCTTAAGTGC ACAGTGCTCT AAACTGA

27

(2) INFORMATION FOR SEQ ID NO: 58:

- 10 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 27 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

20

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 58:

CACGGTAATT AGTGTGATCT GACGAAG

27

25 (2) INFORMATION FOR SEQ ID NO: 59:

- 30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 22 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

35

(iv) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 59:

CGTGCCGCCT TCGTTTCTCT TT

22

(2) INFORMATION FOR SEQ ID NO: 60:

- 45 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 22 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

55

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 60:

TTCGCTTCGG GGTGGATCTG TG

22

5

(2) INFORMATION FOR SEQ ID NO: 61:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 28 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

15

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

20

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 61:

CAAAACTGAC TTACGAGTCA CGTTTGAG

28

25

(2) INFORMATION FOR SEQ ID NO: 62:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 25 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

30

(ii) MOLECULE TYPE: cDNA

35

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 62:

GATGTATGCT TCGTTATTCC ACGCC

25

45

(2) INFORMATION FOR SEQ ID NO: 63:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 21 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

50

(ii) MOLECULE TYPE: cDNA

55

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:

GGTCAAACCT CCAGGGACGC C

21

5 (2) INFORMATION FOR SEQ ID NO: 64:

- (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 24 base pairs
(B) TYPE: nucleic acid
10 (C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

15 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 64:

GCGGTAATGT GTGAAAGCGT TGCC

24

(2) INFORMATION FOR SEQ ID NO: 65:

- 25 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 18 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

30 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

35 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 65:

40 TCCCTTGTGG CCTGTGTG

18

(2) INFORMATION FOR SEQ ID NO: 66:

- 45 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 19 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 66:

TCCTTCATCG GCTCTTCGA

19

5 (2) INFORMATION FOR SEQ ID NO: 67:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 18 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

15 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 67:

GATGCCAAGG CATCCACC

18

25 (2) INFORMATION FOR SEQ ID NO: 68:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 19 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

30 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 68:

CCTCCCCACGT CCTTCATCG

19

40 (2) INFORMATION FOR SEQ ID NO: 69:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 22 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

45 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 69:

AAGGTTCTG ACTAGGTTGG GC

22

(2) INFORMATION FOR SEQ ID NO: 70:

5

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 22 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

10

- (ii) MOLECULE TYPE: cDNA

- (iii) HYPOTHETICAL: NO

15

- (iii) ANTI-SENSE: NO

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 70:

20

GGTGAAGTGC TTGCATGGAT CT

22

(2) INFORMATION FOR SEQ ID NO: 71:

25

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 23 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: cDNA

30

- (iii) HYPOTHETICAL: NO

- (iii) ANTI-SENSE: NO

35

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 71:

ACCTGTGAGT TTTCGTTCTT CTC

23

40

(2) INFORMATION FOR SEQ ID NO: 72:

45

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 24 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: cDNA

- (iii) HYPOTHETICAL: NO

50

- (iii) ANTI-SENSE: NO

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 72:

55

CTATTTGTTG AGTTTGAGA GGTT

24

(2) INFORMATION FOR SEQ ID NO: 73:

- 5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 23 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 73:
 ATTTTCCGTA TCACCGATGA TAC

23

20 (2) INFORMATION FOR SEQ ID NO: 74:

- 25 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 22 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

30 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 74:
 ACGAAGTAAA GGTGTGTTTT CT

22

40 (2) INFORMATION FOR SEQ ID NO: 75:

- 45 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 25 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

50 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 75:
 GAGAGGTTAC TCTCTTTTAT GTCAG

25

(2) INFORMATION FOR SEQ ID NO: 76:

- 5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 275 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

 (iii) HYPOTHETICAL: NO

15 (iii) ANTI-SENSE: NO

15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 76:

AAGGAGCACC ACGAAAACGC CCCAACTGGT GGGCGTAGG CCGTGAGGGG TTCTTGTCTG	60
TAGTGGGCGA GAGCCGGGTG CATGACAACA AAGTTGCCA CCAACACACT GTTGGGTCT	120
GAGGCAACAC TCGGACTTGT TCCAGGTGTT GTCCACCGC CTTGGTGGTG GGGTGTGGTG	180
TTTGAGAACT GGATAGTGGT TGCGAGCATC AATGGATAACG CTGCCGGCTA GCGGTGGCGT	240
GTTCTTTGTG CAATATTCTT TGGTTTTGT TGTGT	275

(2) INFORMATION FOR SEQ ID NO: 77:

- 30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 278 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

 (iii) HYPOTHETICAL: NO

 (iii) ANTI-SENSE: NO

40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 77:

AAGGAGCACC ACGAAAAGCA CCCAACTGG TGGGTGCGA GCCGTGAGGG GTTCCCGTCT	60
GTAATGGACG GGGGCCGGNT GCGAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGTCCGTC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TGGTCTTCGT GGCCGGCGTT	240
CATCGAAATG TGTAATTCT TCCTTAACTC TTGTGTGT	278

(2) INFORMATION FOR SEQ ID NO: 78:

- 55 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 278 base pairs
 (B) TYPE: nucleic acid

(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 78:

AAGGAGCACC ACGAAAAGCA CCCCAACTGG TGGGTGCGA GCCGTGAGGG GTTCCCGTCT	60
GTAGTGGACG GGGGCCGGGT GCGAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGTCCGTC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TGGTCTTCGT GGCCGGCGTT	240
CATCGAAATG TGTAATTCT TTTTTAACTC TTGTGTGT	278

20

(2) INFORMATION FOR SEQ ID NO: 79:

25

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 280 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

30

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 79:

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGTGCGA GCCGTGAGGG GTTCCCGTCT	60
GTAGTGGACG GGGGCCGGNT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TAGTCCTTG GTGTGATGCG	240
CTCGTCGAAA TGTGTAATTG CTTCTTGTT GTNTGTGT	280

40

45

(2) INFORMATION FOR SEQ ID NO: 80:

50

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 281 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

55

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 80:

10	AAGGAGCACC ACGAAAAGCA TCCAATTGG TGGGTGCGA GCCGTGAGGG GTTCTCGTCT	60
	GTAGTGGACG AAAACCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
	GAGACAACAC TCGGTCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
15	TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCG TAGTCCTTG TGGCTGATGC	240
	GTTCATCAAA ATGTGTAATT TCTTTTTGG TTTNTGTGTG T	281

(2) INFORMATION FOR SEQ ID NO: 81:

20 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 280 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

25 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 81:

35	AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGTGCGA GCCGTGAGGG GTTCCCCTCT	60
	GTAGTGGACG GGGGCCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
	GAGACAACAC TCGGTCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
40	TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TAGCCCTTGC GGCTGATGCG	240
	TTCGNCGAAA TGTGTAATT CTTCTCTGGT TTCTGTGTGT	280

(2) INFORMATION FOR SEQ ID NO: 82:

45 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 282 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 82:

5	AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCGA GCCGTGAGGG GTTCTCGTCT	60
	GTAGTGGACG GNAGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
	GAGACAACAC TCGGTCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
10	TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCG TAGTCCTTCG TGGCTGATGC	240
	GTTCATCGAA ATGTGTAATT TCTTCTTTGG TTTTGGGTGT GT	282

(2) INFORMATION FOR SEQ ID NO: 83:

- | | |
|----|-------------------------------|
| 15 | (i) SEQUENCE CHARACTERISTICS: |
| | (A) LENGTH: 282 base pairs |
| | (B) TYPE: nucleic acid |
| | (C) STRANDEDNESS: single |
| | (D) TOPOLOGY: linear |
| 20 | (ii) MOLECULE TYPE: cDNA |
| | (iii) HYPOTHETICAL: NO |
| 25 | (iii) ANTI-SENSE: NO |

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 83:

30	AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCGA GCCGTGAGGG GTTCCCGTCT	60
	GTAGTGGACG GGGGCCGGGT GCACAAACAGC AAATGATCGC CAGACACACT ATTGGGCCCT	120
	GAGACAACAC TCGGTCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
35	TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TAGTCCTTCG GGGCTGATGT	240
	GTTCATCAA AATGTGTAAT TTCTTTTNG GTTTTNGTGT GT	282

(2) INFORMATION FOR SEQ ID NO: 84:

- | | |
|----|-------------------------------|
| 40 | (i) SEQUENCE CHARACTERISTICS: |
| | (A) LENGTH: 281 base pairs |
| | (B) TYPE: nucleic acid |
| | (C) STRANDEDNESS: single |
| | (D) TOPOLOGY: linear |
| 45 | (ii) MOLECULE TYPE: cDNA |
| | (iii) HYPOTHETICAL: NO |
| 50 | (iii) ANTI-SENSE: NO |

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 84:

55	AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCGA GCCGTGAGGG GTTCTCGTCT	60
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5 GTAGTGGACG GGAGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT 120
 GAGACAAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT 180
 TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCG TAGTCCTTCG TGGCTGATGC 240
 GTTCATTGAA ATGTGTAATT TCTTCTCTGG TTTTGTGTG T 281

10 (2) INFORMATION FOR SEQ ID NO: 85:

15 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 280 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iv) ANTI-SENSE: NO

25 (xii) SEQUENCE DESCRIPTION: SEQ ID NO: 85:

25 AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGTGCGA GCCGTGAGGG GTTCCCGTCT 60
 GTAGTGGACG GGGGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT 120
 GAGACAAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT 180
 TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TAGTCCTTGT GGCTGATGCG 240
 CTCGTCGAAA TGTGTAATT TCTCTTGGT TTTTGTGTGT 280

35 (2) INFORMATION FOR SEQ ID NO: 86:

40 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 282 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

45 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iv) ANTI-SENSE: NO

50 (xiii) SEQUENCE DESCRIPTION: SEQ ID NO: 86:

50 AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGTGCGA GCCGTGAGGG GTTCCCGTCT 60
 GTAGTGGACG GGGGCCGGGT GCGAACACAGC AAATGATTGC CAGACACACT ATTGGGCCCT 120
 GAGACAAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTTGGTGT 180
 TTGAGTATTG GATAGTGGTT GCGAGCATCT AGATGAGCGC GTAGTCCTTG TGGCTGATGC 240

GTTCGTCGAA ATGTGTAATT TCTTCTTTGG GTTTTGTGT GT

282

5 (2) INFORMATION FOR SEQ ID NO: 87:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 281 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

15 (iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 87:

AAGGAGCACC ACGAAAAGCA CCCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG GNAGCCGGNT GCGCAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGNCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTNGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGGCGCG TAGTCCTTG TGACTGATGC	240
GTTCATCAAA ATGTGTAATT TCTTTTTGN NTTTNGTGTG T	281

30 (2) INFORMATION FOR SEQ ID NO: 88:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 281 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

40 (iv) ANTI-SENSE: NO

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 88:

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG GGAACCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGTGCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TAGTCCTTG TGGCTGACGC	240
GTTCATCGAA ATGTGTAATT TCTTCTTTGG TTTTTGTGTG T	281

55 (2) INFORMATION FOR SEQ ID NO: 89:

5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 280 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

10 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89:

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCAGA GCCGTGANGG GTTCCCGTCT	60
GTAGTGGACG GGGGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
20 GAGACAACAC TCGGTCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAGCGCA TAGTCCTTAG GGCTGATGCG	240
TTCGTCGNAA TGTGTAATT CTTCTTGGT TTTTGTGTGT	280

25 (2) INFORMATION FOR SEQ ID NO: 90:

30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 282 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

35 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 90:

AAGGAGCACC ACGAAAAGCA TCCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG AAAACCGGGT GCACAAACAGC AAATAATTGC CAGACACACT ATTGGGCCCT	120
45 GAGACAACAC TCGGTCGATC CGTGTGGTGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TAGTCCTTCG TGGCTGACGT	240
GTTCATCGAA ATGTGTAATT TCTTNNTTA ACTCTTGTGT GT	282

50 (2) INFORMATION FOR SEQ ID NO: 91:

55 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 280 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

5 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 91:

AAGGAGCACC ACGAAAAGCA CCCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG GGAGCCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
15 GAGACAACAC TCGGTCAGTC CGTGTGGTGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TAGTCCTTGT GACTGACGTG	240
20 TTCATCGAAA TGTGTAATTCTTTCTAAC TCTTGTGTGT	280

(2) INFORMATION FOR SEQ ID NO: 92:

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 281 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

30 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 92:

AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG AAAGCCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
40 GAGACAACAC TCGGTCGAAC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TGGTCTTCAT GGCCGGCGTG	240
45 TTCATCGAAA TGTGTAATAT CTTCTCTGGT TTTCGGTGTG T	281

(2) INFORMATION FOR SEQ ID NO: 93:

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 280 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

55 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 93:

AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG AAAACCGGNT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGTCGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TGGTCTTCAT GGCCGGCGTG	240
TTCATCGAAA TGTGTAATT TTCTTNNAC TCTTGTGTGT	280

10

(2) INFORMATION FOR SEQ ID NO: 94:

15

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 280 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

20

(ii) MOLECULE TYPE: cDNA

25

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 94:

AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG AAAGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGTCGAAC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TGGTCTTCAT GGCCGGCGTG	240
TTCATCGAAA TGTGTAATT TTCTTGGT TTTNGTGTGT	280

35

(2) INFORMATION FOR SEQ ID NO: 95:

40

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 281 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

45

(ii) MOLECULE TYPE: cDNA

50

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 95:

	AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
5	GTAGTGGACG AAAACCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
	GAGACAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
	TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TAGTCCTCG NGGNNCNGCGT	240
10	GTTCATCGAA ATGTGTAATT TCTNTNTAA CTCTNGTGTG T	281

(2) INFORMATION FOR SEQ ID NO: 96:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 281 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- 20 (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: NO

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 96:

	AAGGAGCACC ACGAAAAGCA TCCCAATTGG TGGGTGTGA GCCGTGAGGG GTTCTCGTCT	60
30	GTAGTGGACG AAAACCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
	GAGACAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
	TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TAGTCCTCG GGGCCGGCGT	240
35	GTTCATCGAA ATGTGTAATT TCTTTTTAA CTCTGTGTG T	281

(2) INFORMATION FOR SEQ ID NO: 97:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 280 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- 45 (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: NO

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 97:

	AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT	60
55	GTAGTGGACG AAAACCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120

5 GAGACAAACAC TCGGTGGAAC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT 180
 TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TGGTCTTCAT GGCCGGCGTG 240
 TTCATCGAAA TGTGTAATT TCTCTTTAAC TCTTGTGTGT 280

(2) INFORMATION FOR SEQ ID NO: 98:

- 10 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 280 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

- 15 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO

- 20 (xii) SEQUENCE DESCRIPTION: SEQ ID NO: 98:

25 AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT 60
 CTAGTGGACG AAAACCGGGT GCACAACAGN AAATGATTGC CAGACACACT ATTGGGCCCT 120
 GAGACAAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT 180
 TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TGGTCTTCAT GGCCNGCGTG 240
 TTCATCCAAA TGTGTAATT TCTTTTTAAC TCTTGTGTGT 280

(2) INFORMATION FOR SEQ ID NO: 99:

- 35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 280 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

- 40 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO

- 45 (xiii) SEQUENCE DESCRIPTION: SEQ ID NO: 99:

50 AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT 60
 CTAGTGGACG AAAACCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT 120
 GAGACAAACAC TCGGTGATC CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT 180
 TGAGTATTGG ATAGTGGTTG CGAGCATCTA GATGAACGCG TGGTCTTCAT GGCCGGCGTG 240
 TTCATCGAAA TGTGTAATT TCTTTTTAAC TCTTGTGTGT 280

(2) INFORMATION FOR SEQ ID NO: 100:

5

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 281 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10

(ii) MOLECULE TYPE: cDNA

15

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 100:

20

AAGGAGCACC ACGAAAAGCA CCCCAACTGG TGGGGTGCAGA GCGGTGAGGG GTCCTCGCCT	60
GTAGTGGCG GGGGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT	120
GAGGCAACAC TCGGCTCGTT CTGAGTGGTG TCCCTCCATC TTGGTGGTGG GGTGTGGTGT	180
TTGAGTATTG GATACTGGTT GCGAGCATCT AAACGGATGC GTGGCCGGCA ACGGTGGCGT	240
GTTCGTTGAA ATGTGTAATT TCTTTTTGG TTTTTGTGTG T	281

25

(2) INFORMATION FOR SEQ ID NO: 101:

30

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 274 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

35

(ii) MOLECULE TYPE: cDNA

40

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 101:

45

AAGGAGCACC ACGAAAAGCA TCCCAACAAG TGGGGTGCAA NCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACC AAAGCCGGGT GCACGACAAC AAGCAAAGCC AGACACACTA TTGGGTCTG	120
AGGCAACACT CGGGCTCTGT TCGAGAGTTG TCCCACCATC TTGGTGGTGG GGTGTGGTGT	180
TTGAGAATTG GATACTGGTT GCGAGCATCA AATGGATGCG TTGCCCTACG GGTAGCGTGT	240
TCTTTTGTGC AATTTTATTC TTTGGTTTT GTGT	274

50

(2) INFORMATION FOR SEQ ID NO: 102:

55

- (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 293 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 102:

AAGGGACACC	ATTTCCCACT	CGATGAAC	GGAAACATAA	AGTAGGCATC	TGTAGTGGAT	60
ATCTACTTGG	TGAATATGTT	TTGTAAATCC	TGTCCACCCC	GTGGATGGGT	AGTCGGCAA	120
ACGTCGGACT	GTCATAAGAA	TTGAAACGCT	GGCACACTGT	TGGGTCCCTGA	GGCAACACGT	180
TGTGTTGTCA	CCCTGCTTGG	TGGTGGGGTG	TGGACTTTGA	CTTCTGAATA	GTGGTTGCGA	240
GCATCTAAC	ATAGCCTCGC	TCGTTTCGA	GTGGGGCTGG	TTTTGCAATT	TTA	293

(2) INFORMATION FOR SEQ ID NO: 103:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 296 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 103:

AAGGAGCACC ATTTCCAGT CGGATGAAC T AGGGAACATA AAGTAGGCAT CTGTAGTGGG	60
TATCTACTTG GTGAATATGT TTTGTAAATC CTGTCCACCC CCGTGGATGG GTAGTCGGCA	120
AAACGTCGGA CTGTCATAAG AATTGAAACG CTGGCACACT GTTGGGTCT GAGGCAACAC	180
GTTGTGTTGT CACCCCTGCTT GGTGGTGGGG TGTGGACTTT GACTTCTGAA TAGTGGTTGC	240
GAGCATCTAA ACATAGCCTC GCTCGTTTC GAGTGAGGCT GGTTTTGCA ATTITA	296

(2) INFORMATION FOR SEO ID NO: 104:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 274 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 104:

AAGGAGCACC ACGAAGAGCA CTCCAATTGG TGGGTGCGA GCCGTGAGGG GTCATCGTCT	60
GTAGTGGACG AAGACCGGGT GCACGACAAC AAGCTAACCC AGACACACTA TTGGGTCCCTG	120
15 AGGCAACACC CTCGGGTGCT GTCCCCCAT CTTGGTGGTG GGGTGTGGTG TTTGAGAAATT	180
GGATAGTGGT TGCGAGCATIC AAAATGTATG CGTTGTCGTT CTCGGCAACG TGTTCTTTT	240
GTGCAATTTA TTCTTGGTT TTTGTAGTGT TTGT	274

20 (2) INFORMATION FOR SEQ ID NO: 105:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 278 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

30 (iii) ANTI-SENSE: NO

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 105:

AAGGAGCACC ACGAAGAGCA CTCCAATTGG TGGGTGCGA GCCGNGAGGG GTCATCGTCT	60
GTAGTGGACG AAGACTGGGT GCACGACAAC AAAGCAAGCC AGACACACTA TTGGGTCCCTG	120
40 AGGCAACACC CTCGGGTGCT GCCCCTCCAT CTTGGTGGTG GGGTGTGGTG TTTGAGAACT	180
GGATAGTGGT TGCGAGCATIC AAAATGTATG CGTTGTCGTT TCGCGACAAC GTGTTCTTTT	240
TGTGCAATTT TAATTCTTTT GGTTTGGTA GTGTTGT	278

45 (2) INFORMATION FOR SEQ ID NO: 106:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 276 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

55 (iii) ANTI-SENSE: NO

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 106:

AAGGAGCACC ACGAGAAGCA CTCCAATTGG TGGGTGCAA GCCGTGAGGG GTCATCGTCT	60
GTAGTGGACG AAGACCCGGT GCACGACAAC AAGCAAAGCC AGACACACTA TTGGGTCTG	120
10 AGGCAACACC CTCGGGTGCT GTCCCCCAT CTTGGTGGTG GGGTGTGGTG TTTGAGAACT	180
GGATAGTGGT TGCGAGCATC AAAATGTATG CGTTGTCGTT CGCGGCAACG TGTTCTTTT	240
GTGCAATTT TATTCTTGG TTTTGTAGT GTTTGT	276

15 (2) INFORMATION FOR SEQ ID NO: 107:

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 277 base pairs
 (B) TYPE: nucleic acid
 20 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

25 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 107:

AAGGAGCACC ACGAAAAGCA CCCCAATTGG TGGGTGCAA GCCGTGAGGG GTTCCCGCCT	60
GTAGTGGCGC GGGCCGGGTG CGAACAGCA AATGATTGCC AGACACACTA TTGGGCCCTG	120
35 AGGCAACACT CGGATCGATT GAGTGCTTGT CCCCCCATCT TGGTGGTGGG GTGTGGTGT	180
TGAGAACTGG ATAGTGGTTG CGAGCATCTA AATGAACGCA CTGCCGATGG TGGTGTGTT	240
GTTTTGTGTA ATTATTATTCT TTGGTTTTG TGTTGT	277

40 (2) INFORMATION FOR SEQ ID NO: 108:

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 282 base pairs
 (B) TYPE: nucleic acid
 45 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

50 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 108:

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGC_G GCCGTNAGGG GTTCTCGTCT 60
 5 GTAGTGGATG GCAGCCGGGT GCACANCAGC AAATGATTGC CAGACACACT ATTGGGCCCT 120
 GAGACAACAC TCGGTCA_GT CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGNGTT 180
 TGAGTATTGG ATAGTGGTTG CGANCATCTA GATGAACGCG TAGTCCTCNG TGGCTGACGT 240
 10 GTTCATCAAA ATGTGTAATT TCTTTTANGG GTTNTGGTGT CT 282

(2) INFORMATION FOR SEQ ID NO: 109:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 280 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

20 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 109:

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGC_G GCCGNAGGG GTTCTCGCCT 60
 30 GTAGTGNCG AGGGCCCGAT GCACAACAAAC ACATGATTGC CAGACACACT ATTGGGCCCT 120
 GANACAACAC TCGGCCAGTC CGTGTGGTGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT 180
 TGAGTATNGG ATAGTN_GTTG NGANCATCTA AACGGCTGCG TN_GNCNN_GAA CGGTGGCGTG 240
 35 TT_GCNTAAAA TGTGTAATT_T CTTTNNNGGT TTGGGTGTNT 280

(2) INFORMATION FOR SEQ ID NO: 110:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 280 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

45 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 110:

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGC_G GCCGTGAGGG GTTCTCGCCT 60
 55 GTAGTGGCG ANGGCCGGGT GCACAACAAAC AAATGATTGC CAGACACACT ATTGGGCCCT 120
 GAGACAACAC TCGGCCAGTC CGTGTGGTGT CCCNCCATCT TGGTGGTGGG GTGTGGTGT 180

TGAGTATTGG ATAGTGGTTG CGAGCATCTA AANGNTGCG TTGCCGNNA CNGTGGCGTN 240

5 TTCGNTAAAA TGTGTAANTT CTTTTTNGGT TTGTGTGTGT 280

(2) INFORMATION FOR SEQ ID NO: 111:

- (i) SEQUENCE CHARACTERISTICS:
 10 (A) LENGTH: 471 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

15 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 111:

ATCGAAGATC CCGGCTTCTT CATAAGCTCC CACACGAATT GCTTGATTCA CTGGTTAGAC 60

25 GATTGGGTCT GTAGCTCAGT TGGTTAGAGC GCACCCCTGA TAAGGGTGAG GTCGGCAGTT 120

CGAACATGCC CAGACCCACC AATTGTTGGT GTGCTGCCTG ATCCGATACG GGGCCATAGC 180

TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGG AGTCGATCC TCCTTGCTC 240

30 CACCATCTAA AACAAATCGTC GAAAGCTCAG AAATGAATGT TCGTGGATGA ACATTGATTT 300

CTGGTCTTTG CACCAGAACT GTTCTTTAAA AATTGGGTA TGTGATAGAA GTAAGACTGA 360

ATGATCTCTT TCACTGGTGA TCATTCAAGT CAAGGTAAAA TTTGCGAGTT CAAGCGCGAA 420

35 TTTTCGGCGA ATGTCGTCTT CACAGTATAA CCAGATTGCT TGGGGTTATA T 471

(2) INFORMATION FOR SEQ ID NO: 112:

- (i) SEQUENCE CHARACTERISTICS:
 40 (A) LENGTH: 520 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

45 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 112:

ATCGAAGACA TCAGCTTCTT CATAAGTATC CACACGAATT GCTTGATTCA TAGTCGAACG 60

55 AATGCTGTAA CGCGACCCGT GTTATAGGTC TGTAGCTCAG TTGGTTAGAG CGCACCCCTG 120

	ATAAGGGTGA GGTGGCAGT TCAAATCTGC CCAGACCTAC CAATTGCTTG GTCGAGAAGA	180
5	ATACGGGCC ATAGCTCAGC TGGGAGAGCG CCTGCCTGAC ACAGCAGGAGG TCAGCGGTT	240
	GATCCCGCTT GGCTCCACCA CTCTCTCGTG TTGCGGTGAG TGTAAAGAG TTCAGAAATG	300
	ATGCCGCTTC AGGTTTGCTC TGTTGAGTGC TGATTCTGG TCTTTGACC GGTACGAAA	360
10	TCGTTCTTA AAAATTGGA TATGTGATAG AAGTGACTGA TTAATTGCTT TCACTGGCAA	420
	TTGATCTGGT CAAGGTAAAA TTTGTAGTTC TCAACACGCA AATTTTCGGC GAATGTCGTC	480
	TTCACGATTG AGACAGTAAC CAGATTGCTT GGGTTATAT	520

15 (2) INFORMATION FOR SEQ ID NO: 113:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 504 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

25 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 113:

30	ATCGAAGACA CCGGCTTCGT CATAAGCTCC CACACGAATT GCTTGATTCA CTTGCGAAAG	60
	GCGATTGGGT TTAGACCCGA GAGTAACGAT TGGGTCTGTA GCTCAGTTGG TTAGAGCGCA	120
35	CCCCTGATAA GGGTGAGGTC GGCAGTCGA ATCTGCCAG ACCCACCAAT CGAAGGGGCC	180
	ATAGCTCAGC TGGGAGAGCG CCTGCTTGC ACGCAGGAGG TCAGCGGTT GATCCCGCTT	240
	GGCTCCACCA TTAACTCTAG TCGCCGAAAG CTCAGAAATG AGTGTGTTACC AGGATGAGGT	300
40	TGATTGCCTG GGTTGAACAT TGATTCTGG ACTTGCGCC AGAACTGTT GTTAAAAAATT	360
	TGGGTATGTG ATAGAAGTAG ACCGATGTGT TGCTTCACT GGCAGCATGT CGCGTCAAGG	420
	TAAAATTGTC GTGTTCTCTA TGCAAATTT CGCGAATGT CGTCTTCACG TTATAGACAG	480
45	TAACCAAGATT GCTTGGGGTT ATAT	504

(2) INFORMATION FOR SEQ ID NO: 114:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 499 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

55 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 114:

	ATCGAAGACT TCAGCTTCTT CATAAGTTCC CACACGAATT GCTTGATTCA CTTGCGAAAA	60
10	GCGATTGGGT TGAGACCCGA GAGTGACGAT TGGGTCTGTA GCTCAGTTGG TTAGAGCGCA	120
	CCCCTGATAA GGGTGAGCTC GGCAGTCGA ATCTGCCAG ACCCACCAAT TGTCGGGATG	180
15	GCCAGTGTCA AATGGGGCCA TAGCTCAGCT GGGAGAGCGC CTGCTTGCA CGCAGGAGGT	240
	CAGGAGTTCG ATCCTCCTTG GCTCCACCAT CAACTCACGA TCGCTGAAAG CTCAGAAATG	300
20	AACATTGGTA GTTCAATGTT GATTTCTGGT CTTTGCAGCA GAACTGTTCT TTAAAAAATTT	360
	GGGTATGTGA TAGAAGTGAC TAACAGCGTG TTTCACTGCA CGTTGTTAAT CAAGGCAAAA	420
25	TTTGCAGTT CAAGCGCGAA TTTTCCGGCA ATGTCGTCTT CACGTTACGA ATCTATAACC	480
	AGATTGCTTG GGGTTATAT	499

25

(2) INFORMATION FOR SEQ ID NO: 115:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 468 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

30

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

35

(iii) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 115:

	ATCGACGACA TCAGCTGTCT CATAAGCTCC CACACGAATT GCTTGATTCA TTGAAGAAGA	60
	CGATTAGGTT AGCAACCTTC GATTGGTCT GTAGCTCAGT TGGTTAGAGC GCACCCCTGA	120
45	TAAGGGTGAG GTCCGGCAGTT CGAACATGCC CAGACCCACC AATTTGCTGG GGCCATAGCT	180
	CAGCTGGGAG AGCGCCTGCC TTGCACGCAG GAGGTCAGCG GTTCGATCCC GCTTGGCTCC	240
	ACCACCCCGC TTGCCAGTT GTCAAAGCTT AGAAATGAAT ATTCCGCTCG AATATTGATT	300
50	TCTGAACCTT ATCAGAACATCG TTCTTTAAAA ATTTGGGTAT GTGATAGAAA GATAGACTGG	360
	ACAGCACTTT CACTGGTGTG TGTCAGGCT AAGGTAAAAT TTGTGAGTAA TTACAAGTTT	420
	TCGGCGAATG TTGTCTTCAC AGTATAACCA GATTGCTTGG GGTTATAT	468

55

(2) INFORMATION FOR SEQ ID NO: 116:

- (i) SEQUENCE CHARACTERISTICS:

 - (A) LENGTH: 246 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 116:

TAAGGAAAAAG GAAACCTGTG AGTTTCGTT CTTCTCTGTT TGTCAGTTT TGAGAGGTAA 60
ATTCTTCTCT ATACTGTTTG TTCTTTGAAA ACTAGATAAG AAAGTTACTA AAGTTACCAT 120
AAATAGGTAAC CTATTATGA CACAAGTAAC CGAGAACATCTGAAAGTGA ATCTTTCATC 180
TGATTGGAAG TATCATCGCT GATACGAAAAA ATCAGAAAAAA CAACCTTTAC TTCATCGAAG 240
TAAATT 246

(2) INFORMATION FOR SEQ ID NO: 117:

- (i) SEQUENCE CHARACTERISTICS:

 - (A) LENGTH: 246 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 117:

CTAAGGAAAA GGAAACCTGT GAGTTTCGTT TCTTCTCTAT TTGTTCAAGTT TTGAGAGGGTT	60
AGTACTTCTC AGTATGTTG TTCTTTGAAA ACTAGATAAG AAAGTTAGTA AAGTTAGCAT	120
AGATAATTAA TTATTTATGA CACAAGTAAC CGAGAACAT CTGAAAGTGA ATCTTTCATC	180
TGATTGGAAG TATCATCGCT GATACGGAAA ATCAGAAAAA CAACCTTTAC TTCGTAGAAG	240
TAAATT	246

(2) INFORMATION FOR SEQ ID NO: 118:

- (i) SEQUENCE CHARACTERISTICS:

 - (A) LENGTH: 246 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

5 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 118:

TAAGGAAAAG GAAACCTGTG AGTTTCGTT CTTCTCTGTT TGTCAGTTTG	60
TTACTTCTCT GTATGTTGT TCTTGAAAAA CTAGATAAGA AAGTTAGTAA AGTTAGCATA	120
15 AGTAGTGTAA CTATTTATGA CACAAGTAAC CGAGAACAT CTGAAAGTGA ATCTTTCATC	180
TAATTCGACG TATCATCGCT GATACAGACA ATTAGAAAAA CAACCTTTAC TTGACCGAAG	240
TAAATT	246

20 (2) INFORMATION FOR SEQ ID NO: 119:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 363 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

25 (ii) MOLECULE TYPE: cDNA

30 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 119:

GGCCTATAGC TCAGCTGGTT AGAGCGCACG CCTGATAAGC GTGAGGTCGA TGGTTCGAGT	60
CCATTTAGGC CCACCTTTTC TTTCTGACAG AAGAAACACT GTATAACCTA TTTAAGGGC	120
40 CTTAGCTCAG CTGGGAGAGC GCCTGCTTTC CACCGAGGAG GTCAGCGGTG CGATCCCGCT	180
AGGCTCCACC AAAATTGTTT TTTGAAAAGT AGATAAGAAA GTTAGTAAAG TTAGCATAAA	240
45 TAGGTAACCA TTTATGACAC AAGTAACCGA GAATCATCTG AAAGTGAATC TTTCATCTGA	300
TTGGAAGTAT CATCGCTGAT ACGAAAAATC AGAAAAACAA CCTTTACTTC ATCGAAGTAA	360
ATT	363

50 (2) INFORMATION FOR SEQ ID NO: 120:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 496 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 120:

TAAGGAAAAG GAAACCTGTG AGTTTCGTT CTTCTCTATT TGTCAGTTT TGAGAGGTTA	60
CTCTCTTTA TGTCAGATAA AGTATGCAAG GCACATGCT TGAAGCATCG CGCCACTACA	120
15 TTTTGACGG GCCTATACCT CAGCTGGTTA GAGCGCACGC CTGATAAGCG TGAGGTCGAT	180
GGTTCGAGTC CATTAGGCC CACTTTCT TTCTGACATA AGAAATACAA ATAATCATAAC	240
20 CCTTTTACGG GGCTTAGCT CAGCTGGGAG AGCCGCTGCT TTGCACGCAG GAGGTCAGCG	300
GTTCGATCCC GCTAGGCTCC ACCAAAATTG TTCTTGAAA ACTAGATAAG AAAGTTAGTA	360
AAGTTAGCAT AGATAATTAA TTATTTATGA CACAAGTAAC CGAGAACATCT GTGAAAGTGA	420
25 ATCTTCATC TGATTGGAAG TATCATCGCT GATAACGGAA ATCAGAAAAA CAACCTTAC	480
TTCGTAGAAG TAAATT	496

(2) INFORMATION FOR SEQ ID NO: 121:

30 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 498 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 121:

TAAGGAAAAG GAAACCTGTN AGTTNCGTN CTTCTCTGTT TGTCAGTTT TNAGAGGTTA	60
45 CTCTCTTTNA TGTCAGATAA AGTACGCACG GCACGTTGCC TTGGGCAAAG AGCCACTACA	120
TTATTGACGG CCCTATAGCT CAGCTGGTTA GAGCGCACGC CTGATAAGCG TGAGGTCGAT	180
GGTTCGAGTC CATTAGGCC CACTTTCT TTCTGACAGA AGAAATCATT TGCACATCCT	240
50 ATTAATAAGG GNCCCTAGCT CAGCTGGGAG AGCCGCTGCT TTGCACGCAG GAGGTCAGCG	300
GTTCGATCCC GCTAGGCTCC ACCAAAATT GTTCTTGAA AACTAGATAA GAAAGTTAGT	360
AAAGTTAGCA TAAGTAGTAT AACTATTTAT GACACAAGTA ACCGAGAACATC ATCTGAAAGT	420
55 GAATCTTCA TCTAATTCCA CGTATCATCG CTGATAACAGA CAATNGAAA AACAACCTT	480

ACTTCGACGA AGTAAATT

498

5 (2) INFORMATION FOR SEQ ID NO: 122:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 229 base pairs
 (B) TYPE: nucleic acid
 10 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

15 (iii) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 122:

TAAGGATAAG GATAACTGTC TTAGGACGGT TTGACTAGGT TGGGCAAGCG TTTTTTTAAT	60
CTTGTATTCT ATTCCCTTTG CATTGTTAACG CGTTGTTCC AAAACATTAA GTTTACGATC	120
AAGTATGTTA TGTAATAAT ATGGTAACAA GTAAATTACAC ATATAATAAT AGACGTTAA	180
GAATATATGT CTTTAGGTGA TGTAACTTG CATGGATCAA TAATTTACA	229

(2) INFORMATION FOR SEQ ID NO: 123:

- 30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 248 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 123:

TAAGGATAAG GAAGAAGCCT GAGAAGGTTT CTGACTAGGT TGGGCAAGCA TTTATATGTA	60
AGAGCAAGCA TTCTATTCA TTTGTGTTGT TAAGAGTAGC GTGGTGAGGA CGAGACATAT	120
AGTTTGTGAT CAAGTATGTT ATTGTAAGA AATAATCATG GTAACAAGTA TATTCACGC	180
50 ATAATAATAG ACGTTTAAGA GTATTTGTCT TTTAGGTGAA GTGCTTGCAT GGATCTATAG	240
AAATTACA	248

(2) INFORMATION FOR SEQ ID NO: 124:

- 55 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 224 base pairs

(ii) MOLECULE TYPE: cDNA

5 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 118:

TAAGGAAAAAG GAAACCTGTG AGTTTCGTT CTTCTCTGTT TGTCAGTTT TGAGAGGTTA	60
TTACTCTCT GTATGTTGT TCTTTGAAAA CTAGATAAGA AAGTTAGTAA AGTTAGCATA	120
15 AGTAGTGTAA CTATTTATGA CACAAGTAAC CGAGAACAT CAT CTGAAAAGTGA ATCTTTCATC	180
TAATTGACG TATCATCGCT GATACAGACA ATTAGAAAAA CAACCTTAC TTGACGAAG	240
TAAATT	246

20 (2) INFORMATION FOR SEQ ID NO: 119:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 363 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

30 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 119:

GGCCTATAGC TCAGCTGGTT AGAGCGCACG CCTGATAAGC GTGAGGTCGA TGGTTCGAGT	60
CCATTTAGGC CCACTTTTTC TTTCTGACAG AAGAAACACT GTATAACCTA TTTAAGGGC	120
40 CTTAGCTCAG CTGGGAGAGC GCCTGCTTGCACCGAGGAG GTCAAGCGGTT CGATCCCCGT	180
AGGCTCCACC AAAATTGPTC TTTGAAAAGT AGATAAGAAA GTTAGTAAAG TTAGCATAAA	240
TAGGTAACCA TTTATGACAC AAGTAACCGA GAATCATCTG AAAGTGAATC TTTCATCTGA	300
45 TTGGAAGTAT CATCGCTGAT ACGAAAAATC AGAAAAACAA CCTTTACTTC ATCGAAGTAA	360
ATT	363

50 (2) INFORMATION FOR SEQ ID NO: 120:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 496 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 120:

TAAGGAAAAG	GAAACCTGTG	AGTTTCGTT	CTTCTCTATT	TGTCAGTTT	TGAGAGGTTA	60
CTCTCTTTA	TGTCAGATAA	AGTATGCAAG	GCACATGCT	TGAAGCATCG	CGCCACTACA	120
15 TTTTGACGG	GCCTATAGCT	CAGCTGGTTA	GAGGCCACGC	CTGATAAGCG	TGAGGTCGAT	180
GGTCGAGTC	CATTTAGGCC	CACTTTTCT	TTCTGACATA	AGAAATACAA	ATAATCATAAC	240
20 CCTTTTACGG	GGCCTTAGCT	CAGCTGGGAG	AGCGCCTGCT	TTGCACGCAG	GAGGTCAGCG	300
GTTCGATCCC	GCTAGGCTCC	ACCAAAATTG	TTCTTGAAA	ACTAGATAAG	AAAGTTAGTA	360
AAGTTAGCAT	AGATAATTAA	TTATTTATGA	CACAAGTAAC	CGAGAACAT	CTGAAAGTGA	420
25 ATCTTCATC	TGATTGGAAG	TATCATCGCT	GATACGGAAA	ATCAGAAAAA	CAACCTTAC	480
TTCGTAGAAG	TAATT					496

(2) INFORMATION FOR SEQ ID NO: 121:

30 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 498 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 121:

TAAGGAAAAG	GAAACCTGTN	AGTTTNCGTN	CTTCTCTGTT	TGTCAGTTT	TNAGAGGTTA	60
45 CTCTCTTNA	TGTCAGATAA	AGTACGCACG	GCACGTTGCC	TTGGGCAAAG	AGCCACTACA	120
TTATTGACGG	GCCTATAGCT	CAGCTGGTTA	GAGGCCACGC	CTGATAAGCG	TGAGGTCGAT	180
GGTCGAGTC	CATTTAGGCC	CACTTTTCT	TTCTGACAGA	AGAAATCATT	TGCACATCCT	240
50 ATTAATAAGG	GNCCCTAGCT	CAGCTGGGAG	AGCGCCTGCT	TTGCACGCAG	GAGGTCAGCG	300
GTTCGATCCC	GCTAGGCTCC	ACCCAAAATT	GTCTTTGAA	AACTAGATAA	GAAAGTTAGT	360
AAAGTTAGCA	TAAGTAGTAT	AACTATTTAT	GACACAAGTA	ACCGAGAAC	ATCTGAAAGT	420
55 GAATCTTCA	TCTAATTGCA	CGTATCATCG	CTGATAACAGA	CAATTNGAAA	AAACACCTT	480

ACTTCGACGA AGTAAATT

498

5 (2) INFORMATION FOR SEQ ID NO: 122:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 229 base pairs
 (B) TYPE: nucleic acid
 10 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

15 (iii) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 122:

TAAGGATAAG GATAACTGTC TTAGGACGGT TTGACTAGGT TGGGCAAGCG	60
CTTGATTTCT ATTCCCTTTG CATTGTTAACG CGTTGTTCC AAAACATTTA GTTTACGATC	120
AAGTATGTTA TGTAATAAT ATGGTAACAA GTAAATTACATATAATAAT AGACGTTAA	180
GAATATATGT CTTTAGGTGA TGTAACTTG CATGGATCAA TAATTTACA	229

25 (2) INFORMATION FOR SEQ ID NO: 123:

- 30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 248 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

40 (iii) ANTI-SENSE: NO

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 123:

TAAGGATAAG GAAGAAGCCT GAGAAGGTTT CTGACTAGGT TGGGCAAGCA	60
TTTATATGTA AGAGCAAGCA TTCTATTTCA TTTGTGTTGT TAAGAGTAGC GTGGTGAGGA CGAGACATAT	120
AGTTTGTGAT CAAGTATGTT ATTGTAAGA AATAATCATG GTAACAAGTA TATTCACGC	180
50 ATAATAATAG ACGTTTAAGA GTATTTGTCT TTTAGGTGAA GTGCTTGCAT GGATCTATAG	240
AAATTACA	248

55 (2) INFORMATION FOR SEQ ID NO: 124:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 224 base pairs

- (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

10 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 124:

15 CAAATGGAGT TTTTATTTT TATTTATCTT AAACACCCAT TAATTTTTC GGTGTTAAAA	60
CCCAAATCAA TGTGGTCT CACAACAAAC ACATTTGGTC AGTTGTATC CAGTTCTGAA	120
AGAATGTTT TGAAACAGTTC TTTCAAAACT GAAAACGACA ATCTTCTAG TTCCAAAAAT	180
20 AAATACAAA GGATCAATAAC AATAAGTTAC TAAGGGCTTA TGGT	224

(2) INFORMATION FOR SEQ ID NO: 125:

- 25 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 252 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

30 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 125:

40 CTAATGAAGT TTTTACTTT TTCTTTCAT CTTAATAAA GATAAATACT AAACAAAACA	60
TCAAAATCCA TTTATTTATC GGTGGTAAAT TAAACCCAAA TCCCTGTTG GTCTCACAC	120
TAACATATTT GGTCAGATTG TATCCAGTTC TGAAAGAACAA TTTCCGCTTC TTTCAAAACT	180
45 GAAAACGACA ATCTTCTAG TTCCAAATAA ATACCAAAGG ATCAATAACAA TAAGTTACTA	240
AGGGCTTATG GT	252

(2) INFORMATION FOR SEQ ID NO: 126:

- 50 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 608 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

55 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 126:

10	AACGAAAGAT TGACGATTGG TAAGAATCCA CAACAAGTTG TTCTTCATAG ATGTATCTGA	60
	GGGTCTGTAG CTCAGTTGGT TAGAGCACAC GCTTGATAAG CGTGGGGTCA CAAGTTCAAG	120
	TCTTGTCAAG CCCACCATGA CTGGACTGG TTGAAGTTAT AGATAAAAAGA TACATGATTG	180
15	ATGATGTAAG CTGGGGACTT AGCTTAGTTG GTAGAGCGCC TGCTTGCAC GCAGGAGGTC	240
	AGGAGTTCGA CTCTCCTAGT CTCCACCAGA ACTTAAGATA AGTTCGGATT ACAGAAATTA	300
	GTAAATAAAAG ATTGAGATCT TGGTTTATTA ACTTCTGTGA TTTCATTATC ACGGTAATT	360
20	GTGTGATCTG ACGAAGACAC ATTAACATCA TAACAGATTG GCAAAATTGA GTCTGAAATA	420
	AATTGTTCAC TCAAGAGTTT AGGTTAACAGA ATTAATCTAG ATGAATTGAG AACTAGCAA	480
	TTAACTGAAT CAAGCGTTT GGTATGTGAA TTAGATTGA AGCTGTACAG TGCTTAAGTG	540
25	CACAGTGCTC TAAACTGAAA TGTGAAGTT ACTAACTTGT AGGTAAACATC GACTGTTGG	600
	GGTTGTAT	608

(2) INFORMATION FOR SEQ ID NO: 127:

30

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 269 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

35

(ii) MOLECULE TYPE: cDNA

40

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 127:

45	AACGAAAGAT TGACGATTGG TAAGAATCCA CGACAAGTTG TTCTTCATAG ATGTATCTGA	60
	GGGTCTGTAG CTCAGTTGGT TAGAGCACAC GCTTGATAAG CGTGGGGTCA CAAGTTCAAG	120
	TCTTGTCAAG CCCACCATGA CTGGACTGG TTGAAGTTAT AGAAAAGAAG ATACATAACT	180
50	GATGATGTAAG GCTGGGGACT TAGCTTAGTT GGTAGAGCGC CTGCTTGCAC CGCACGGAGGT	240
	CAGGAGTTCG ACTCTCCTAG TCTCCACCA	269

55

(2) INFORMATION FOR SEQ ID NO: 128:

- (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 249 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

5

- (ii) MOLECULE TYPE: cDNA

10

- (iii) ANTI-SENSE: NO

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 128:

15

AACGAAAGAT TGATGGCCGG TAAGAATCCA CAACAAGTTG TTCTTCGAAG ATGTATCTGA
GGGTCTGTAG CTCAGTTGGT TAGAGCACAC GCTTGATAAG CGTGGGTCA CAAGTTCAAG
TCTTGTCAAG CCCACCAAAT CTGAAAGATA TGTCGTTCAT TATGATTAAG GCTGGGGACT
TAGCTTAGTT GGTAGAGCGC CTGCTTGCA CGCAGGAGGT CAGGAGTTCG ACTCTCCTAG
TCTCCACCA

35

- (2) INFORMATION FOR SEQ ID NO: 129:

30

- (i) SEQUENCE CHARACTERISTICS:

 - (A) LENGTH: 283 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

35

(iii) ANTI-SENSE: NO

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 129:

40

AACGAAAGAT TCACGATTGG TAAGAATCCA CAACAAGTTG TTCTTCATGA CGATGTATCT
GAGGGTCTGT AGCTCAGTTG GTTAGAGCAC ACGCTTGATA AGCGTGGGGT CACAAGTTCA
AGTCTTGTCA GACCCACCAA ATCTGACTAA CAAGCATTAT TAAATGCTGA ATACAGAAAA
ACAGAGACAT TGACTTATTG ATAAGCTGGG GACTTAGCTT AGTTGGTAGA GCGCCTGCTT
TGCACGCCAGG AGGTCAAGGAG TTTCGACTCTC CTAGTCTCCA CCA

28

- (2) INFORMATION FOR SEQ ID NO: 130:

55

- (i) SEQUENCE CHARACTERISTICS:

 - (A) LENGTH: 283 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 130:

AACGAAAGAT TGGTGACCGG TAAGAATCCA CAACAAGTTG TTCTTCGAAG ATGTATCTGA	60
GGGTCTGTAG CTCAGTTGGT TAGACCACAC GCTTGATAAG CGTGGGGTCA CAAGTTCAAG	120
TCTTGTCAAG CCCACCACTA CTGACGAAGT GATGAATAAT CACAAGCTGC TAGATGAAAA	180
GATATGTCGT TCATTATGAT TAAAGCTGGG GACTTAGCTT AGTTGGTAGA GCGCCTGCTT	240
TGCACGCAGG AGGTCAGGAG TTCGACTCTC CTAGTCTCCA CCA	283

20 (2) INFORMATION FOR SEQ ID NO: 131:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 808 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

25 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

30 (iii) ANTI-SENSE: NO

35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 131:

TAAGGAAGAT CGAGAACATTGG AAAGAGGTCG GATTTATCCG GATGATCCTT CTCCATCTTA	60
TTAGAACATA GATCGCAGGC CAGTCAGCCT GACCGATCGCT TGCAGGCCGT CGGCCTTCGT	120
40 TTCTCTTCT TCATTGTTGA TTGCTCACGG GCCGTACCGC AGCTGACGCT GCTGGCCCTG	180
CGCAGGCGCG GCCCATCAGG GCCGACGGCC GGTCGGCCTT GCNAAGCTTC GCTTCGGGTT	240
GGATCTGTGG ATCGCGTAGT AGCGTTGCG TCGGTATCTG GGCTTGTAGC TCAGTTGGTT	300
45 AGAGCACACG CTTGATAAGC GTGGGGTCGG AGGTTCAAGT CCTCCCAGGC CCACCAAAGTT	360
ACTTGATGAG GGGCCGTAGC TCAGCTGGA GAGCACCTGC TTTGCAAGCA GGGGGTCGTC	420
GGTTCGATCC CGTCCGGCTC CACCATCATG TTGGTGTGA GACGGATATT GGCAATCAAC	480
50 AAAAGAAAAGA AACAAAGTTG CGGACTNTTA CGAAAGTCTG CCTGTTCTGT ATGAAATCGT	540
GAAGAGAAGA TGTAATCGGA TCAACTGAAG AGTTGATGTC GCAAGAAGCT TGCTCAAGCC	600
TTGCATAATG ATTGATGTGT TTAACCGCCA TCACCGATTG TATCTCGAGA AGCTGGTCTT	660
55 TCTGCTGATA CTGTTGAAAC GAGCATTGAGC AGTCGAATGG CAACATTGG CGTCGCATAA	720

TGCGGCTTA AGAGCTGAGT TTTGATGGAT ATTGGCAATG AGAGTGATCA AGTGTCTAA	780
5 GGGCATTGGT GGATGCCTTG GCATGCAC	808

(2) INFORMATION FOR SEQ ID NO: 132:

- 10 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 808 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 132:

TAAGGAGGAT CGAGAACATTGG AAAGAGGCCG GATTTATCCG GATGATCCTT CTCCATCTTA	60
25 TTAGAACATA GATCGCAGNC CAGTCAGCCT GACGATCGCT TGCAGGCCGT CGGCCTTCGT	120
TTCTCTTTCT TCATTGTTGA TTGCTCACGG GCCGTACCGC AGCTGACGCT GCTGGCCCTG	180
CGCAGGCCG GNCCATCAGG GCCGACGGCC GGTCGGCCTT GCGAAGCTTC GCTTCGGGT	240
30 GGATCTGTGG ATCGCGTAGT AGCGTTGCG TCGGTATCTG GGCTTGTAGC TCAGTTGGTT	300
AGAGCACACG CTTGATAAGC GTGGGGTCGG AGGTTCAAGT CCTCCCAGGC CCACCAAAGTT	360
ACTTGATGAG GGGCCGTAGC TCAGCTGGGA GACCACCTGC TTTGCAAGCA GGGGGTCGTC	420
35 CGTTCGATCC CGTCCGGCTC CACCATCATG TTGGTGTGA GACGGATATT GGCAATCAAC	480
AAAAGAAAGA AACAAAGTTG CGGACTNTTA CGAAAGTCTG CCTGTTCTGT ATCAAATCGT	540
GAAGAGAAGA TGTAATCGGA TCAACTGAAG AGTTGATGTC GCAAGAAGCT TGCTCAAGCC	600
40 TTGCATAATG ATTGATGTGT TTAACCGCCA TCACCGATTG TATCTCGAGA AGCTGGTCTT	660
TCTGCTGATA CTGTTGAAAC GAGCATTGTC AGTCGAATGG CAACATTGG CGTCGCATAA	720
45 TGCAGCTTTA AGAGCTGAGT TTTGATGGAT ATTGGCAATG AGAGTGATCA AGTGTCTAA	780
GGGCATTGGT GGATGCCTTG GCATGCAC	808

(2) INFORMATION FOR SEQ ID NO: 133:

- 50 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 353 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

55 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 133:

10	CCTTAAAGAA CTGTTCTTTG CAGTGCTCAC ACAGATTGTC TGATGAAAAG TAAATAGCAA	60
	GGCGTCTTGC GAAGCAGACT GATACTCCC CTTCGTCTAG AGGCCAGGA CACCGCCCTT	120
	TCACGGCGGT AACAGGGGTT CGAATCCCCT AGGGGACGCC ACTTGCGCGG TAATGTGTGA	180
15	AACCGTTGCC ATCACTATCT CAAAACTGAC TTACGAGTCA CGTTTGAGAT ATTTGCTCTT	240
	TAAAAATCTG GATCAAGCTG AAAATTGAAA CACAGAACAA CGAAAGTTGT TCGTGAGTCT	300
	CTCAAATTT CGCAACACGA TGATGAATCG TAAGAAACAT CTTGGGTTG TGA	353

20 (2) INFORMATION FOR SEQ ID NO: 134:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 515 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

30

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 134:

35	CCTTAAAGAA CTGTTCTTTG CAGTGCTCAC ACAGATTGTC TGATGAAAAA CGAGCAGTAA	60
	AACCTCTACA GGCTTGTAGC TCAGGTGGTT AGAGCGCACC CCTGATAAGG GTGAGGTCGG	120
40	TGGTTCAAGT CCACTCAGGC CTACCAAATT TTCCCTGAAT ACTGCGTTGT GAAATAACTC	180
	ACATACTGAT GTATGCTTCG TTATTCCACG CCTTGTCTCA GGAAAATTAA TCGGTAAAGA	240
	GGTTCTGACT ACACGATGGG GCTATAGCTC AGCTGGGAGA GCGCCTGCTT TGCACCCAGG	300
45	AGGTCTGCGG TTCGATCCCG CATAGCTCCA CCATATCGTG AGTGTITACG AAAAAAATCT	360
	TCAGAGTGTAA CCTGAAAGGG TTCACTGCGA AGTTTGCTC TTTAAAATC TGGATCAAGC	420
	TGAAAATTGA AACACAGAAC AACGAAAGTT GTTCGTGAGT CTCTCAAATT TTGCAACAC	480
50	GATGATGAAT CGTAAGAAC ATCTTCGGGT TGTGA	515

(2) INFORMATION FOR SEQ ID NO: 135:

(i) SEQUENCE CHARACTERISTICS:

55

- (A) LENGTH: 353 base pairs
- (B) TYPE: nucleic acid

(C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 135:

CCTTAAAGAA GCGTACTTTG CAGTGCTCAC ACAGATTGTC TGATGAAAAG TAAATAGCAA	60
GGCGTCTTGC GAAGCAGACT GATACTCCC CTTCGTCTAG AGGCCAGGA CACCGCCCTT	120
TCACGGCGGT AACAGGGGTT CGAATCCCCT AGGGGACGCC ACTTGCGCGG TAATGTGTGA	180
AAGCGTTGCC ATCAGTATCT CAAAAGTAC TTACCGAGTCA CGTTTGAGAT ATTTGCTCTT	240
TAAAAATCTG GATCAAGCTG AAAATTGAAA CACAGAACAA CGAAAGTTGT TCGTGAGTCT	300
CTCAAATTTT CGCAACACGA TGATGAATCG TAAGAAACAT CTTCGGGTTG TGA	353

25 (2) INFORMATION FOR SEQ ID NO: 136:

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 481 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

30 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

35 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 136:

CCTTAAAGAA CTGTTCTTGT AAGTGCTCAC ACAGATTGTC TGATGAAAAA CGAGCAGTAA	60
AACCTCTACA GGCTTGAGC TCAGGGTGT AGAGCGCACC CCTGATAAGG GTGAGGTCGG	120
TGGTTCAAGT CCACTCAGGC CTACCAAATT TTCCCTGAAT ACTGCGTTGT GAAATAACTC	180
ACATACTGAT GTATGCTTCG TTATTCCACG CCTTGTCTCA GGAAAAATTG TCGGTAAAGA	240
GGTTCTGACT ACACGATGGG GCTATAGCTC AGCTGGGAGA GCGCCTGCTT TGCACGCAGG	300
AGGTCTGCGG TTCGATCCCG CATAGCTCCA CCATCTCGTG AGTGTGTTACG AAAAAAATCT	360
TCAGAGTGTA CCTGAAAGGG TTCACTGCGA AGTTTGCTC TTTAAAAATC TGGATCAAGC	420
TGAAAATTGA AACACAGAAC AACGAAAGTT GTTCGTGAGT CTCTCAAATT TTGCAACAC	480
G	481

(2) INFORMATION FOR SEQ ID NO: 137:

5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 392 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

10 (iii) HYPOTHETICAL: NO

15 (iii) ANTI-SENSE: NO

15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 137:

CCTTAAAGAA	CGGTACTTTG	AAGTGCTCAC	ACAGATTGTC	TGATGAAAAG	TGAATAGCAA	60
GGCGTCTTGC	GATTGAGACT	TCAGTGTC	CTTCGTCTAG	AGGCCAGGA	CACCGCC	120
TCACGGCGGT	AACAGGGTT	CGAATCCCCT	AGGGGACGCC	AGCGTCAAA	CTGATGAGGT	180
CAAACCTCCA	GGGACGCCAC	TTGCTGGTTT	GTGAGTGAAA	GTCACCTGCC	TTAATATCTC	240
AAAACTGACT	TACGAGTCAC	TTTGAGATA	TTTGCTCTTT	AAAAATCTGG	ATCAAGCTGA	300
AAATTGAAAC	ACAGAACAAAC	GAAAGTTGTT	CGTGAGTCTC	TCAAATTTTC	GCAACACGAT	360
GATGAATCGT	AAGAACATC	TTCGGGTTGT	GA			392

30 (2) INFORMATION FOR SEQ ID NO: 138:

35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 515 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA

40 (iii) HYPOTHETICAL: NO

45 (iii) ANTI-SENSE: NO

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 138:

CCTTAAAGAA	ACGGTCTTTG	AAGTGCTCAC	ACAGATTGTC	TGATGAAAAA	CGAGCAGTAA	60
AACCTCTACA	GGCTTGTAGC	TCAGGTGGTT	AGAGCGCACC	CCTGATAAGG	CTGAGGTCGG	120
TGGTTCAAGT	CCACTCAGGC	CTACCAAATT	TTCCCTGAAT	ACTGCGTTGT	GAAATAACTC	180
ACATACTGAT	GTATGCTTCG	TTATTCCACG	CCTTGTCTCA	GGAAAAATT	TCGGTAAAGA	240
GGTTCTGACT	ACACGATGGG	GCTATAGCTC	AGCTGGGAGA	GCGCCTGCTT	TGCACGGCAGG	300
AGGTCTGCGG	TTCGATCCCCG	CATAGCTCCA	CCATCTCGTG	AGTGTTCAGG	AAAAAATACT	360

TCAGAGTGTA CCTGAAAGGG TTCACTGCAG AGTTTGCTC TTTAAAAATC TGGATCAAGC 420
 5 TGAAAATTGA AACACAGAAC AACGAAAGTT GTTCGTGAGT CTCTCAAATT TTGCAACAC 480
 GATGATGAAT CGTAAGAAC ATCTTCGGGT TGTGA 515

(2) INFORMATION FOR SEQ ID NO: 139:

- 10 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 365 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 139:

25 CTAAGGATAT ATTCCGAACA TCTTCTTCGG AAGATGCGGA ATAACGTGAC ATATTGTATT 60
 CAGTTTGAA TGTTTATTAA ACATTCAAAT ATTTTTGGT TAAAGTGATA TTGCTTTGA 120
 AAATAAAGCA GTATGCGAGC GCTTGACTAA AAAAATTGT ACATTGAAAA CTAGATAAGT 180
 30 AAGTAAAATA TAGATTTAC CAAGCAAAAC CGAGTGAATA AAGAGTTTA AATAAGCTTG 240
 AATTCTATAAG AAATAATCGC TAGTGTTCGA AAGAACACTC ACAAGATTAA TAACGCCATT 300
 AAATCTTTT ATAAAAGAAC GTAACCTCAT GTTAACGTTT GACTTATAAAA AATGGTGGAA 360
 35 ACATA 365

(2) INFORMATION FOR SEQ ID NO: 140:

- 40 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 548 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

45 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 140:

CTAAGGATAT ATTCCGAACA TCTTCTTCAG AAGATGCGGA ATAACGTGAC ATATTGTATT 60
 55 CAGTTTGAA TGTTTATTAA ACATTCAAAT ATTTTTGGT TAAAGTGATA TTGCTTATGC 120

	GAGCNCTTGA CAATCTATT CTTTTAAAGA AAGCGGTTGT CAGACAATGC ATTAAGAAAA	180
5	ATTAAGCGG AGTTTACTTT TGTAAATGAG CATTTGATTT TTTGAAAATA AAGCAGTATG	240
	CGAGCGCTTG ACTAAAAGA AATTGTACAT TGAAAACGT ATAAGTAAGT AAAATATAGA	300
	TTTTACCAAG CAAAACCGAG TGAATAAAGA GTTTTAAATA AGCTTGAATT CATAAGAAAT	360
10	AATCGCTAGT GTTCGAAAGA ACACTCACAA GATTAATAAC CGCTTAAAT CTTTTATAA	420
	AAGAAAACGT TTAGCAGACA ATGAGTTAA TTATTTAAA GCAGAGTTA CTTATGTAAA	480
	TGAGCATTTA AAATAATGAA AACGAAGCCG TATGTGAGCA TTTGACTTAT AAAAATGGTG	540
15	GAAACATA	548

(2) INFORMATION FOR SEQ ID NO: 141:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 471 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: cDNA

- 25 (iii) HYPOTHETICAL: NO

- (iv) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 141:

	CTAAGGATAT ATTCTGGAACA TCTTCTTCAG AAGATGCGGA ATAACGTGAC ATATTGTATT	60
35	CAGTTTGAA TGTAAATTTA ACATTCAAAT ATTTTTGGT TAAAGTGATA TTGCTTATGC	120
	CGAGCGCTTGA CAATCTATT CTTTTAAAGA AAGCGGTTGT CAGACAATGC ATTAAGAAAA	180
	ATTAAGCGG AGTTTACTTT TGTAAATGAG CATTTGATTT TTTGAAAATA AAGCAGTATG	240
40	CGAGCGCTTG ACTAAAANGA AATTGTACAT TGAAAACGT ATAAGTAAGT AAAATATAGA	300
	TTTTACCAAG CAAAACCGAG TGAATAAAGA GTTTGAAATA AGCTTGAATT CATAAGAAAT	360
	AATCGCTAGT GTTCGAAAGA ACACTCACAA GATTAATAAC CGCTTAAAT CTTTTATAA	420
45	AAGAACGTAA CTTCATGTTA ACGTTGACT TATAAAAATG GTGGAAACAT A	471

(2) INFORMATION FOR SEQ ID NO: 142:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 383 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: cDNA

- 55 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 142:

	CTAAGGATAT ATTCGGAACA TCTTCAG AAGATGCGGA ATAACGTGAC ATATTGTATT	60
10	CAGNTTGAA TGTTTATTAA ACATTCAAAA AATGGGCCTA TAGCTCAGCT GGTTAGAGCG	120
	CACGCCCTGAT AACCGTGAGG TCGGTGGTTC GAGTCCACTT AGGCCACCA TTATTTGTAC	180
	ATTGAAAACT AGATAAGTAA GTAAAATATA GATTTTACCA AGCAAAACCG AGTGAATAAA	240
15	GAGTTTTAAA TAAGCTTGAA TTCATAAGAA ATAATCGCTA GTGTTGAAA GAACACTCAC	300
	AAGATTAATA ACGCGTTAA ATCTTTTAT AAAAGAACGT AACTTCATGT TAACGTTGA	360
	CTTATAAAAA TGGTGGAAAC ATA	383

20

(2) INFORMATION FOR SEQ ID NO: 143:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 351 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

30

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 143:

	CTAAGGATAT ATTCGGAACA TCTTCYTCAAG AAGATGCGGA ATAATGTGAC ATATTGTATT	60
	CAGTTTGAA TGTTTATTAA ACATTCAAAT ATTTTTGGT TAAAGTGATA TTGCTTATGC	120
40	GAGCGCTTGA CTAAAAAGAA ATTGTACATT GAAAAGTAGA TAAGTAAGTA AAANTATAGA	180
	TTTTACCAAG CAAAACCGAG TGAATAAGA GTTTAAATA AGCTGAATT CATAAGAAAT	240
	AATCGCTAGT GTTCGAAAGA ACACTCACAA GATTAATAAC GCGTTAAAT CTTTTATAAA	300
45	AAGAACGTAA CTTCATGTTA ACGTTTGACT TATAAAAATG GTGGAAACAT A	351

45

(2) INFORMATION FOR SEQ ID NO: 144:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 263 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

55

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 144:

10	CTAAGGATAT ATT CGGAACA TCTTCTACGA AGATGAGGGA ATAACGTGAC ATATTGTATT	60
	CAGTTTGAA TGTTTATTAA CATTCAATTG TACATTGAAA ACTAGATAAG TAAGTAAGAT	120
	TTTACCAAGC AAAACCGAGT GAATAGAGTT TTAAATAAGC TTGAATTCAT AAATAATCGC	180
15	TAGTGTCGA AAGACNTCCA CAAGATTAAT AACTAGTTT AGCTATTTAT TTTGAATAAC	240
	AATTCAAAAT ATGGTGGGAC ATA	263

(2) INFORMATION FOR SEQ ID NO: 145:

20	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 247 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

25	(ii) MOLECULE TYPE: cDNA
----	--------------------------

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 145:

35	AAGGATAAGG AACTGCACAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGTG GGGCCTTAGC	60
	TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGC GGTTCGATCC CGCTAGGCTC	120
	CATTGGTGAG AGATCACCAA GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA	180
40	ACAAGAAAAT AAACCGAAAA CGCTGTAGTA TTAATAAAGA GTTTATGACT GAAAGGTCAA	240
	AAAATAA	247

(2) INFORMATION FOR SEQ ID NO: 146:

45	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 375 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

50	(ii) MOLECULE TYPE: cDNA
----	--------------------------

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 146:

5	AAGGAAATGG AACACGTTA TCGTCTTATT TAGTTTGAG AGGTCTTGTG GGGCCTTAGC	60
	TCAGCTGGGA GAGCGCCTGC TTNGCACGCA GGAGGTCAGC GGTCGATCC CGCTAGGCTC	120
	CATCAGGATA CANTCCTACT AAACTTAATA CAAGTGAAGT TGAACACGCA ACTCACTTCC	180
10	TAGAAAATA GACAATCTTC GCTTGTGTGC AAGGCACACA TGGTCAGATT CCTAATTTC	240
	TACAGAACTT TCGCTAAAGC GAGCGTTGCT TAGTATCCTA TATAATAGTC CATNGAAAAT	300
	TGAATATCTA TATCAAATTG CACGATCTAG AAATAGATTG TGGAAACGTA ACAAGAAATT	360
15	AACCCGNAAA CGCTG	375

(2) INFORMATION FOR SEQ ID NO: 147:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 244 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

- (iii) HYPOTHETICAL: NO
- (iii) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 147:

35	AAGGATAAGG AACTGCACAT TGGTCTTGTGTT TAGTCTTGAG AGGTCTTGTG GGGCCTTAGC	60
	TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGC GGTCGATCC CGCTAGGCTC	120
	CATTGGTGAG AGATCACCAA GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA	180
	ACAAGAAAAT AAACCGAACAC GCTGTAGTAT TAAAAGAGTT TATGACTGAA AGGTAGAAA	240
40	ATAA	244

(2) INFORMATION FOR SEQ ID NO: 148:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 284 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

- (iii) HYPOTHETICAL: NO
- (iii) ANTI-SENSE: NO

55

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 148:

5	CTAAGGATAT ATTCGGAACA TCTTCTTACG AAGATGCAGG AATAACATTG ACATATTGTA	60
	TTCAGNTGTG AATGCTCATT GGAGNATTCA TNGCATNATT TGGTNCATTG ACANCTAGAT	120
	AAGNAAGTAA AATTTATGAT TTTACCAAGC AAAACCGAGT GAATTAGAGT TNTNNAACAA	180
10	GCTTGATTT CAAAAAGAAA TAATCGCTAG TGTTGAAAG AACACTCACA GATTANTAAC	240
	ATCTTGGTT TTCACCCGAC TTGTTCGTNT CGAAAGTCAA AAAA	284

(2) INFORMATION FOR SEQ ID NO: 149:

- 15 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 246 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 149:

30	AAGGATAAGG AACTGCGCAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGAG GGGCCTTAGC	60
	TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGC GGTCGATCC CGCTAGGCTC	120
	CATTGGTGAG AGATCACCAA GTAATGCACA TTGAAAATTG AATATCTATA TCCTATAGTA	180
35	ACAAGAAAAT AAACCGAAAA CGCTGTAGTA TTAATAAGAG TTTATGACTG AAAGGTCAAA	240
	AAATAA	246

(2) INFORMATION FOR SEQ ID NO: 150:

- 40 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 247 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

45 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 150:

55	AAGGATAAGG AACTGCGCAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGAG GGGCCTTAGC	60
	TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGC GGTCGATCC CGCTAGGCTC	120

CATTGGTGAG AGATCACCAA GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA	180
5 ACAAGAAAAT AAACCGAAAA CGCTGTAGTA TTAATAAGAG TTTATGACTG AAAGGTCAGA	240
AAAATAA	247

(2) INFORMATION FOR SEQ ID NO: 151:

- 10 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 247 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

- 15 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO

20
 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 151:

AAGGAAAAGG AACTGCGCAT TGGTCTTGTGTT TAGTCTTGAG AGGTCTTGTG GGGCCTTAGC	60
TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGC GGTCGATCC CGCTAGGCTC	120
CATTGGTGAG AGATCACCAA GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA	180
30 ACAAGAAAAT AAACCGAAAA CGCTGTAGTA TTAATAAGAG TTTATGACTG AAAGGTCAGA	240
AAAATAA	247

(2) INFORMATION FOR SEQ ID NO: 152:

- 35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 244 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

- 40 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO

45
 45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 152:

50 AAGGATAAGG AACTGCGCAT TGGTCTTGTGTT TAGTCTTGAG AGGTCTTGTG GGGCCTTAGC	60
TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGC GGTCGATCC CGCTAGGCTC	120
CATTGGTGAG AGATCACCAA GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA	180
55 ACAAGAAAAT AAACCGAAAC GCTGTAGTAT TAAAAGAGTT TATGACTGAA AGGTCAGAAA	240

ATAA

244

(2) INFORMATION FOR SEQ ID NO: 153:

- 5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 243 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

15 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 153:

20	AAGGATAAGG AACTGCGCAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGTG GGGCCTTAGC	60
	TCAGCTGGGA GAGCGCCTGC TTTGCACGCA GGAGGTCAGC GGTTCGATCC CGCTAGGCTC	120
	CATTGGTGAG AGATCACCAA GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA	180
25	ACAAGAAAAT AAACCGAAAC GCTGTAGTAT TAAAAGAGTT TATGACTGAA AGGTCAAAAA	240
	TAA	243

(2) INFORMATION FOR SEQ ID NO: 154:

- 30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 809 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

40 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 154:

45	TAAGGAAGAT CGAGAACATTG AAAGAGGTCG GATTTATCCG GATGATCCTT CTCCATCTTA	60
	TTAGAACATA GATCGCAGGC CAGTCAGCCT GACGATCGCT TGCAGGCCGT CGGCCTTCGT	120
	TTCTCTTTCT TCATTGTTGA TTGCTCACGG GCCGTACCGC AGCTGACGCT GCTGGCCCTG	180
50	CGCAGGCGCG GCCCATCAGG GCCGAACGGC CGGTGGCCT TGCNAAGCTT CGCTTCGGGG	240
	TGGATCTGTG GATCGCGTAG TAGCGTTGC GTCGGTATCT GGGCTTGTAG CTCAGTTGGT	300
	TAGAGCACAC GCTTGATAAG CGTGGGGTCG GAGGTTCAAG TCCTCCCAGG CCCACCAAGT	360
55	TACTTGATGA GGGGCCGTAG CTCAGCTGGG AGAGCACCTG CTTTGCAAGC AGGGGGTCGT	420

	CGGTTCGATC CCGTCCGGCT CCACCATCAT GTTGGTGTTC AGACGGATAT TGGCAATCAA	480
5	CAAAGAAAG AAACAAGTTT GCGGACTNTT ACGAAAGTCT GCCTGTTCTG TATGAAATCG	540
	TGAAGAGAAG ATGTAATCGG ATCAACTGAA GAGTTGATGT CGCAAGAAC TTGCTCAAGC	600
	CTTGCATAAT GATTGATGTG TTTAACCGCC ATCACCGATT GTATCTCGAG AAGCTGGTCT	660
10	TTCTGCTGAT ACTGTTGAAA CGAGCATTG CAGTCGAATG GCAACATTG GCGTCGCATA	720
	ATGCGGCTTT AAGAGCTGAG TTTTGATGGA TATTGGCAAT GAGAGTGATC AAGTGTCTTA	780
	AGGGCATTGG TGGATGCCTT GGCGATGCAC	809

15 (2) INFORMATION FOR SEQ ID NO: 155:

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 23 base pairs
 (B) TYPE: nucleic acid
 20 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA
 25 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 155:

TGGGGTGAAG TCGTAACAAG GTA

23

(2) INFORMATION FOR SEQ ID NO: 156:

35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 21 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO

45

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 156:

50 CCTTTCCCTC ACGGTACTGG T

21

(2) INFORMATION FOR SEQ ID NO: 157:

55 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 277 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

5 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 157:

AAGGAGCACC ACGAGAAACA CTCCAATTGG TGGGGTGTAA GCCGTGAGGG GTTCTCGTCT	60
15 GTAGTGGACG GAAGCCGGGT GCACAACAAC AAGCAAGCCA GACACACTAT TGGGTCCCTGA	120
GCACAACATCT CTGTTGGTTT CGGGATGTTG TCCCACCATC TTGGTGGTGG GGTGTGGTGT	180
20 TTGAGAATTG GATA GTGGTT GCGAGCATCA ATTGGATGCG CTGCCTTTG GTGGCGTGT	240
CTGTTGTGCA ATTTTATTCT TTGGTTTTG TGTTTAT	277.

(2) INFORMATION FOR SEQ ID NO: 158:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 286 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

30

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 158:

AAGGAGCACC ACGAGAAACA CCCCAATTGG TGGGGTGTGA GCCGTGAGGG GTTCTCGTCT	60
40 GTAGTGGACG AGGGCCGGGT GCACAACAAC AGGCAATCGC CGGACACACT ATTGGGCCCT	120
GAGACAACAC TCGGCCGACT GAGGTCGACG TGGTGTCCCT CCATCTTGGT GGTGGGGTGT	180
45 GGTGTTGAG CATTGAATAG TGGTTGCGAG CATCTAGCCG GATGCGTTCC CCAGTGGTGC	240
GCGTCGTCA AAAATGTGTA ATTTTCTTT TCGTTTTGT GTTCGT	286

(2) INFORMATION FOR SEQ ID NO: 159:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 286 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

55

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 159:

10	AAGGAGCACC ACGAGAAACA CCCCAATTGG TGGGGTGTGA GCCGTGAGGG GTTCTCGTCT	60
	GTAGTGGACG AGGGCCGGGT GCACAAACAC AGGCAATCGC CGGACACACT ATTGGGCCCT	120
	GAGACAACAC TCGGCCGACT GAGGTCGACG TGGTGTCCCT CCATCTTGTT GGTGGGGTGT	180
15	GGTGTGAG CATTGAATAG TGGTGCAG CATCTAGACG GATGCGTTCC CCAGTGGTGC	240
	GGCTTCGTCA AAAATGTGTA ATTTTCTTT TGGTTTTGT GTTCGT	286

(2) INFORMATION FOR SEQ ID NO: 160:

20 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 279 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

25 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 160:

35	AAGGAGCACC ACGAGAAACA CCCCAATTGG TGGGGTGTGA GCCGTGAGGG GTTCTCGTCT	60
	GTAGTGGACG AGGCAGGTAC ACAACGCCA ATCGCCGGAC ACACATTGG GCCTGAGACA	120
	ACACTCGGCC GACTGAGGTC GACGTGGTGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT	180
40	TGAGCATTGA ATAGTGGTTG CGAGCATCTA GCCGGATGCG TTCCCCAGTG GTGCGCGTTC	240
	GTCAAAAATG TGTAATTTT CTTGGTTT TGTGTTCGT	279

(2) INFORMATION FOR SEQ ID NO: 161:

45 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 288 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 161:

5	AAGGAGCACC ACGAGAAACA CTCCAATTGG TGGAGTGTGA GCCGTGAGGG GTTCTCGTCT	60
	GTAGTGGACG AGGGCCGGGT GCACAAACAGC AGACAATCGC CAGACACACT ATTGGGCCCT	120
	GAGACAAACAC TCGGCCGACT TTGGTCGACG TGGTGTCCCT CCATCTTGGT GGTGGGGTGT	180
10	GGTGTGTTGAG CATTGAATAG TGGTTGCGAG CATCTAGACG GATGCGTTGC CCTCGGGCCG	240
	CGTGTTCGTC AAAAATGTGT AATTTTTCT TTTGGTTTT GTGTTCGT	288

(2) INFORMATION FOR SEQ ID NO: 162:

15	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 289 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

20	(ii) MOLECULE TYPE: cDNA
	(iii) HYPOTHETICAL: NO
25	(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 162:

30	AAGGAGCACC ACGAGAAACA CTCCAATTGG TGGAGTGTGA GCCGTGAGGG GTTCTCGTCT	60
	GTAGTGGACG GGAGCCGGGT GCACAAACAAC AGGCAATCGC CAGACACACT ATTGGGCCCT	120
	GAGACAAACAC TCGGCCGGCT TTGAGTCGAA CTGGTGTCCC TCCATCTTGG TGGTGGGGTGT	180
35	TGGTGTGTTGA GCATTGAATA GTGGTTGCGA GCATCTAGAC GGATGCGTTG CCTTCGGGCC	240
	CGCTGTTCGT AAAAATGTG TAATTTTTC TTTGGTTTT TGTGTTCGT	289

(2) INFORMATION FOR SEQ ID NO: 163:

40	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 232 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

45	(ii) MOLECULE TYPE: cDNA
	(iii) HYPOTHETICAL: NO
50	(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 163:

55	AGGGAGCACC GAAACGCATC CCGCGTGGGG TGTGGGTTCG CCCGTGTGTC GCCTCGGGCCG	60
----	--	----

AGGTGTTGGG CAGCAGGCAG TAACCCCGGA ACACTGTTGG GTTTGAGAA CACCCGTGCT	120
GGTGGTGTGC TCCCCGTGGT GCGGGGTGTG GTGTTGAGT GTGGATAGT GGTTGCGAGC	180
5 ATCTGGCAA GACTGTGGTA AGCGGTTTT GTTGATGTT TCTGGTGTG GT	232

(2) INFORMATION FOR SEQ ID NO: 164:

- 10 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 279 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

20

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 164:

AAGGAGCACCC ACGAGAAACA CTCCAATTGG TGGGGTGTGA GCCGTGAGGG GTTCTCGTCT	60
25 GTAGTGGACG AGGGCGGGTG CACAACAACA GCAATGCCA GACACACTAT TGGCCCTGAG	120
ACAAACACTCG GCCGACTTGG TTGAAGTGGT GTCCCTCCAT CTTGGTGGTG GGGTGTGGTG	180
TTTGAGTATT GGATAGTGGT TGCGAGCATC TAATGAACGC GTCGCCGCAA CGGTTACGTG	240
30 TTCGTTTGT GTAATTTTC TATTGGTTTT TGTGTTCGT	279

(2) INFORMATION FOR SEQ ID NO: 165:

- 35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 285 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

45

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 165:

AAGGAGCACCC ACGAGAAACA CTCCAATTGG TGGGGTGTGA GCCGTGAGGG GTTCTCGTCT	60
50 GTAGTGGACG AGGGCCGGGT GCACAACAAC AGGCAATCGC CAGACACACT ATTGGCCCTG	120
AGACAACACT CGGCCGACTT TGGTCGAAGT GGTGTCCCCC CATCTTGGTG GTGGGGTGTG	180
GTGTTGAGT ATTGGATAGT GGTTGCGAAC ATCTAAATGA ACGCGTTGCC GGCAACGGTT	240
55 ACGTGTTCGT TTTAGTGTAA TTTTCTAAT GGTTTTGTG TTCGT	285

(2) INFORMATION FOR SEQ ID NO: 166:

- 5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 384 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

 (iii) HYPOTHETICAL: NO

15 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 166:

AAGGAGCACC ACGAGACCTG GGCGGGCCCC GCAGATCGCG GGATCAGCTG AGCTTCAGG	60
CGATTCTGTG GATGGCCTCG CACCTGTAGT GGGTGGGGGT CTGGTGCACT CAACAAACTT	120
GGCGTGGGAT CGGGAAAGC ATCTGCGAA AATCATCAGA CACACTATTG GGCTTGAGA	180
CAACAGGCC CGAGCCTGCC CGTTGGGGC AGGGGTGTGT TGTTGCCTCA CTTTGGTGGT	240
GGGGGTGGTG TTTGATTGT GGATAGTGGT TGCGAGCATC TAGCGCGCAG AATGTGTGGT	300
CTCACTCCTT GTGGGTGGGG CCTGGTTTG TGTGCGATTG ATGTGCAATT TCTTTGAAA	360
30 CTCATTTTTG GGTTTTGTG TTGT	384

(2) INFORMATION FOR SEQ ID NO: 167:

- 35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 295 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA

 (iii) HYPOTHETICAL: NO

45 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 167:

AAGGAGCACC ACGAAAAACT CCCAATTGG TGGGTGAA GCCGTGAGGG GTTCCCGTCT	60
50 GTAGTGGACG CGGGCCGGGT GCGAACAGC AAGCGAACAG CCGGACACAC TATTGGTCC	120
TGAGGCACACA CTCGGTTTG TCCCCCTCAG GGATTTCTG GGTGTTGTCC CACCATCTG	180
GTGGTGGGGT GTGGTCTTG AGAATTGGAT AGTGGTTGCG AGCATCAAAT GGATGCGTTG	240
55 CCCCTACGGG TAGCGTGTTC TTTTGTCAA TTTTATTCTT GGTTTTGTG TTTGT	295

(2) INFORMATION FOR SEQ ID NO: 168:

- 5 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 279 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

10 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

15

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 168:

AAGGAGCACC ACGAGAACCA CTCCAACCTGG TGGGGTGCAA GCCGTGAGGG GTTCTCGTCT	60
GTAGTGGACG AGAGCCGGGT GCGCGACAAC GAACGAGCCA GACACACTAT TGGGTCCCTGA	120
GGCAACACTC GGGCTTGCCC AGAGCTGTTG TCCCACCATC TTGGTGGTGG GGTGTGGTGT	180
TTGAGAATTG GATACTGGTT GCGAGCATCA AATGGATGCG TTGCCCCCTAC GGGTGGCGTG	240
25 TTCTTTGTG CAATTTATT CTTGGTTTT TGTGTTTGT	279

(2) INFORMATION FOR SEQ ID NO: 169:

- 30 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 286 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 169:

AAGGAGCACC ACGAAAAACA CCCCAACCTGG TGGGGTGTAACGCCGTGAGGG GCTCCCGTCT	60
GTAGTAGACG GGCGCCCCGT CGCGAACAGC AAGCGAGCCA GACACACTAT TGGGTCCCTGA	120
GGCAACACTC GGGCTTGTC TGGACTCGTC CAAGAGTGTGTT GTCACCACAT CTTGGTGGTGT	180
GGGTGTGGTG TTTGAGAATT GGATAGTGGT TGCGAGCATC ACTGGATGCG TTGCCCCCAG	240
50 GGGTAGCGTG TTCTTTGTG CAATTTATTC TGGTTTTGT GTTAGT	286

(2) INFORMATION FOR SEQ ID NO: 170:

- 55 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 265 base pairs
 (B) TYPE: nucleic acid

(C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 170:

AAGGAGCACC ACGAAAAACA CTCCGCATCC GGTGGGGTGT GAGCCGTGAG GGAGCCCGTG	60
CCTGTAGTGG GTGTGGGTTG GGTGCGCGAC AACAAATGGG AAAAATCGCT GGGCACACTA	120
TTGGGCTTTG AGGCAACACC TGGTTTGTTC TGGGTGGTGT CGCTCCATCT TGGTGGTGGG	180
GTGTGGTGTGTT TGAGTTGTGG ATAGTGGTTG CGAGCATCTA AGCAAAAGCT GTTGTGTTGAC	240
20 GGTTTTGTC GAGTGTGTTG TGTGT	265

(2) INFORMATION FOR SEQ ID NO: 171:

25 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 299 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

30

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 171:

AAGGAGCACC ACGAAAAACA CTCCAATTGG TGGGGTGTAA GCCGTGAGGG GTTCTCATCT	60
40 GTAGTGGACG AGAGCCGGGT GCACAAACAGC AAATGAATCG CCAGACACAC TGTTGGGTCC	120
TGAGGCAACA CTCAGGCTTG TCCCATGTTG GGCTTGATCG GGTGCTGTCC CCCCCATCTTG	180
GTGGTGGGGT GTGGTGTGTTG AGTATTGGAT AGTGGTTGCG AGCATCTAAA TGGATACGTT	240
45 GCCAGTAATG GTGGCGTATT CATTGAAAAT GTGTAATTTC CTTCTTTGGT TTTGTGTGT	299

(2) INFORMATION FOR SEQ ID NO: 172:

50 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 299 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

55

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 172:

10	AAGGAGCACC ACGAAAAACA CTCCAATTGG TGGGGTGTAA GCCGTGAGGG GTTCTCATCT	60
	GTAGTGGACG AGAGCCGGGT GCACAAACAGC AAATGAATCG CCAGACACAC TGTTGGGTCC	120
	TGAGGCAACA CTCAGGCTTG TCCCATGTTG GGCTTGATCG GGTGCTGTCC CCCCATCTTG	180
15	GTGGTGGGGT GTGGTGTGGT AGTATTGGAT AGTGGTTGCG AGCATCTAAA TGGATACGTT	240
	GCCAGTAATG GTGGCGTGTGTT CATTGAAAAT GTGTAATTTC CTTCTTTGGT TTTGTGTGT	299

(2) INFORMATION FOR SEQ ID NO: 173:

20	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 298 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

25	(ii) MOLECULE TYPE: cDNA
----	--------------------------

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 173:

35	AAGGAGCACC ACGAAAAACA CTCCAATTGG TGGGGTGTAA GCCGTGAGGG GTTCTCATCT	60
	GTAGTGGACG AGAGCCGGGT GCACAAACAGC AAATGAATCG CCAGACACAC TGTTGGGTCC	120
	TGAGGCAACA CTCAGGCTTG TCCCATGTTG GGCTTGATCG GGTGCTGTCC CCCCATCTTG	180
40	GTGGTGGGGT GTGGTGTGGT AGTATTGGAT AGTGGTTGCG AGCATCTAAA TGGAACGTTG	240
	CCAGTAATGG TGGCGTGTTC ATTGAAAATG TGTAATTTC TTCTTTGGT TTGTGTGT	298

(2) INFORMATION FOR SEQ ID NO: 174:

45	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 300 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

50	(ii) MOLECULE TYPE: cDNA
----	--------------------------

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

55

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 174:

5	AAGGAGCACC ATTTCTCAGT CGAATGAAC T GAGAACATAA AGCGAGTATC TGTAGTGGAT	60
	ACATGCTTGG TGAATATGTT TTATAAATCC TGTCCACCCC GTGGATAGGT AGTCGGCAA	120
	ACGTCGGACT GTCATAAGAA TTGAAACGCT GGCACACTGT TGGGTCCCTGA GGCAACACAT	180
10	TGTGTTGTCA CCCTGCTTGG TGGTGGGGTG TGGTCCTGA CTTATGGATA GTGGTTGCGA	240
	GCATCTAACAC ATAGCCTCGC TCGTTTCGA GTGAGGCTGG TTTTGCAAT TTTATTAGCT	300

(2) INFORMATION FOR SEQ ID NO: 175:

15	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 22 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

20	(ii) MOLECULE TYPE: cDNA
	(iii) HYPOTHETICAL: NO

25	(iii) ANTI-SENSE: NO
----	----------------------

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 175:

30	GGTTTCGGGA TGTTGTCCCCA CC	22
----	---------------------------	----

(2) INFORMATION FOR SEQ ID NO: 176:

35	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 21 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

40	(ii) MOLECULE TYPE: cDNA
	(iii) HYPOTHETICAL: NO
	(iii) ANTI-SENSE: NO

45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 176:
	CGACTGAGGT CGACGTGGTG T

21

50	(2) INFORMATION FOR SEQ ID NO: 177:
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55	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 27 base pairs
	(B) TYPE: nucleic acid
	(C) STRANDEDNESS: single
	(D) TOPOLOGY: linear

5 (iii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 177:

GGTGTGAG CATTGAATAG TGGTTGC

27

(2) INFORMATION FOR SEQ ID NO: 178:

15 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 21 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

25 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 178:

30 GTTGGGCAGC AGGCAGTAAC C

21

(2) INFORMATION FOR SEQ ID NO: 179:

35 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 20 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

40 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 179:

CCGGCAACGG TTACGTGTT

20

50 (2) INFORMATION FOR SEQ ID NO: 180:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 21 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 180:

TCGTTGGATG GCCTCGCACC T

21

(2) INFORMATION FOR SEQ ID NO: 181:

15 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 21 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 181:

ACTTGGCCGTG GGATGCCGGGA A

21

30 (2) INFORMATION FOR SEQ ID NO: 182:

35 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 24 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

40 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 182:

CCCTCAGGGA TTTTCTGGGT GTTG

24

(2) INFORMATION FOR SEQ ID NO: 183:

50 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 23 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

55 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 183:

10 GGACTCGTCC AAGAGTGTG TCC

23

(2) INFORMATION FOR SEQ ID NO: 184:

15 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 21 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

25 (iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 184:

TCGGGCTTGG CCAGAGCTGT T

21

30 (2) INFORMATION FOR SEQ ID NO: 185:

35 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 20 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

45 (iii) ANTI-SENSE: NO

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 185:

50 GGGTGCGCAA CAGCAAGCGA

20

(2) INFORMATION FOR SEQ ID NO: 186:

55 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 19 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 186:

GATGCGTTGC CCCTACGGG

10

19

(2) INFORMATION FOR SEQ ID NO: 187:

15

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 24 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

20

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 187:

CCCTACGGGT AGCGTGTTCT TTTG

24

30

(2) INFORMATION FOR SEQ ID NO: 188:

35

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 23 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

40

(iii) HYPOTHETICAL: NO

45

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 188:

CGGATCGATT GAGTGCTTGT CCC

23

50

(2) INFORMATION FOR SEQ ID NO: 189:

55

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 23 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

55

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 189:

TCTAAATGAA CGCACTGCCG ATG

23

10

(2) INFORMATION FOR SEQ ID NO: 190:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 20 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

15

(ii) MOLECULE TYPE: cDNA

20

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

25

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 190:

TGAGGGAGCC CGTGCCTGTA

20

(2) INFORMATION FOR SEQ ID NO: 191:

30

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 22 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

35

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 191:

45

CATGTTGGGC TTGATCGGGT GC

22

(2) INFORMATION FOR SEQ ID NO: 192:

50

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 20 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

55

(iii) HYPOTHETICAL: NO

5 (iii) ANTI-SENSE: NO

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 192:

CCTGGGTTTG ACATGCACAG

20

10 (2) INFORMATION FOR SEQ ID NO: 193:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 21 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

20 (iii) ANTI-SENSE: NO

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 193:

GCGTAGTACG GTTTCGGTCG G

21

(2) INFORMATION FOR SEQ ID NO: 194:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 21 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

35 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 194:

CGCAAGAAC TTGCTCAAGC C

21

45 (2) INFORMATION FOR SEQ ID NO: 195:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 470 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

55 (iii) ANTI-SENSE: NO

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 195:

	CCTAATGATA TTGATTGCGC TGAAGTGCTC ACACAGATTG TCTGATGAAA AAGTAACGAG	60
10	CAGAAATACC TTTATAGGCT TGTAGCTAG GTGGTTAGAG CGCACCCCTG ATAAGGGTGA	120
	GGTCGGTGGT TCAAGTCCAC TCAGGCCTAC CACTTCTCGA AGTGGAAAAG GTACTGCACG	180
15	TGACTGTATG GGGCTATAGC TCAGCTGGCA GAGCGCCTGC CTTGCACCCA GGAGGTCAGC	240
	GGTCGATCC CGCTTAGCTC CACCATATAG TCCTGTATTT CAATACTTCA GAGTGTACTG	300
	GCAACAGTAT GCTGCGAAGT ATTTTGCTCT TTAACAATCT GGAACAAGCT GAAAATTGAA	360
20	ACATGACAGC TGAAACTTAT CCCTCCGTAG AAGTATTGGG GTAAGGATTA ACCTGTCTA	420
	GAGTCTCTCA AATGTAGCAG CACGAAAGTG GAAACACCTT CGGGTTGTGA	470

(2) INFORMATION FOR SEQ ID NO: 196:

- 25 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 453 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear
- 30 (ii) MOLECULE TYPE: cDNA
 (iii) HYPOTHETICAL: NO
 (iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 196:

	CCTAATGATA TTGATTGCGC TGAAGTGCTC ACACAGATTG TTTGATAGAA ACGTAATGAG	60
40	CAAAAGCGCT ACCTGTTGAT GTAATGAGTC ACTGACTCAT GCTGATACGA ACCGATTAAG	120
	ACAGTCAGTT TAATCGGATT TTCGTGTCCT CATCGTCTAG AGGCCTAGGA CACTGCCCTT	180
	TCACGGCTGT AACAGGGTTT CGAATCCCCT TGGGGACGCC ATTGATAAT GAGTGAAAGA	240
45	CATTATCACC GGTTCTTGGA ACCGAAAACA TCTTAAAGAT GACTCTTGCG AGTCGTGTTT	300
	AAGATATTGC TCTTTAACAA TCTGGAACAA GCTGAAAATT GAAACATGAC ACCTGAAACT	360
	TATCCCTCCG TAGAAGTATT CGGGTAAGGA TTAACCTGTC ATAGAGTCTC TCAAATGTAG	420
50	CAGCACGAAA GTGAAACAC CTTCGGGTTG TGA	453

(2) INFORMATION FOR SEQ ID NO: 197:

- 55 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 248 base pairs
 (B) TYPE: nucleic acid

- (C) STRANDEDNESS: single
(D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 197:

TAAGGATAAG	GAAGAAGCCT	GAGAAGGTTT	CTGACTAGGT	TGGGCAAGCA	TTTATATGTA	60
15 AGAGCAAGCA	TTCTATTCA	TTTGTGTTGT	TAAGAGTAGC	GCGGTCACCA	CGAGACATAT	120
AGTTTGAT	CAAGTATGTT	ATTGAAAGA	AATAATCATG	GTAACAAAGTA	TATTCACGC	180
ATAATAATAG	ACGTTTAAGA	GTATTGCT	TTTAGGTGAA	GTGCTTGCAT	GGATCTATAG	240
20 AAATTACA						248

(2) INFORMATION FOR SEQ ID NO: 198:

- 25 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 23 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

30 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

35

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 198:

GGAAAAGGTA	CTGCACGTGA	CTG	23
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40 (2) INFORMATION FOR SEQ ID NO: 199:

- 45 (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 23 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

50 (iii) ANTI-SENSE: NO

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 199:

GACAGCTGAA ACTTATCCCT CCG

23

(2) INFORMATION FOR SEQ ID NO: 200:

5

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

10

(ii) MOLECULE TYPE: cDNA

15

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 200:

GCTACCTGTT GATGTAATGA GTCAC

25

(2) INFORMATION FOR SEQ ID NO: 201:

20

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 22 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

25

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

30

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 201:

35

GAGTAGCGCG GTGAGGACGA GA

22

(2) INFORMATION FOR SEQ ID NO: 202:

40

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

45

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

50

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 202:

CTTTTATGTC AGATAAAGTA TGCAG

25

55

(2) INFORMATION FOR SEQ ID NO: 203:

- (i) SEQUENCE CHARACTERISTICS:

- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

10 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 206:

15 TTGACTGCA AATGCTCG

18

(2) INFORMATION FOR SEQ ID NO: 207:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

25 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 207:

TCTTAAAGCC GCATTATGC

19

(2) INFORMATION FOR SEQ ID NO: 208:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

40 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

45 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 208:

50 CCTAATGATA TTGATTGCG

20

(2) INFORMATION FOR SEQ ID NO: 209:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 22 base pairs
 - (B) TYPE: nucleic acid

(C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

5 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 209:

ATGACAGGTT ATTCCTTACC CC

22

15

(2) INFORMATION FOR SEQ ID NO: 210:

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 25 base pairs
 (B) TYPE: nucleic acid
 20 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

25

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 210:

GGTGTGGTCC TTGACTTATG GATAG

25

(2) INFORMATION FOR SEQ ID NO: 211:

35

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 21 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

40

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

45

(iii) ANTI-SENSE: NO

50

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 211:

TGGGGCCCG TGTCGTCAA A

21

(2) INFORMATION FOR SEQ ID NO: 212:

55

(i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 24 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

5

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 212:

CGTTTCATA AGCGATCGCA CGTT

24

15

(2) INFORMATION FOR SEQ ID NO: 213:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 235 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single

20

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

25

(iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 213:

30

TAAGGATAAG GAAACCTGTG AATCTTTTC CCTTCTTTG TTCAGTTTG AGAGGTTCAT

60

CTCTCAAAAC GTGTTCTTG AAAACTAGAT AAGAAAAGTT AGTGTAAAAA GACGAAGAGA

120

35

AACCGTAGGT TTTTCTCAA CCAAAACCGA GAATCAAACC GAGAAAGAAT CTTCCTGTT

180

TCATAAGCGA TCGCACGTT ATGAAAACAC AACAACACCT TCGTAAGAAG GATGA

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(2) INFORMATION FOR SEQ ID NO: 214:

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(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 475 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

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(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iii) ANTI-SENSE: NO

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 214:

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TAAGGATAAG GAAACCTGTG AATCTTTTC CCTTCTTTG TTCAGTTTG AGAGGTCAT

60

GACGCTCATA CTGAGTACCA GGTGACACGT TTTGAGGTG TCTCTCGTA TGAGGGCCT

120

ATAGCTCAGC TGGTTAGAGC GCACGCCGA TAAGCGTGAG GTCGGTGGTT CGAGTCCACT 180
 5 TAGGCCACT TTTTGAAATA AACCTTCCTT TTTTATATGT TAATAAGGGG CCTTAGCTCA 240
 GCTGGGAGAG CGCCTGCTT GCACGCAGGA GGTCAAGGGT TCGATCCGC TAGGCTCCAC 300
 CAAAGATAGT TTGTTCTTG AAAACTAGAT AAGAAAAGTT AGTGTAAAAA GACGAAGAGA 360
 10 AACCGTAGGT TTTCTTCAA CCAAAACCGA GAATCAAACC GAGAAAGAAT CTTCCGTT 420
 TCATAAGCGA TCGCACGTT ATGAAAACAC AACAAACACCT TCGTAAGAAG GATGA 475

(2) INFORMATION FOR SEQ ID NO: 215:

- 15 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 463 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

20 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

25 (iii) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 215:

30 TAAGGATAAG GAAACCTGTG AATCTTTTC CCTCTTTTG TTCAGTTTG AGAGGTCAAT 60
 GACGCTCATA CTGAGTACCA GGTGACACGT TTTTGAGGTG TCTCTTCGTA TGAGGGCCT 120
 ATAGCTCAGC TGGTTAGAGC GCACGCCGA TAAGCGTGAG GTCGGTGGTT CGAGTCCACT 180
 35 TAGGCCACT TTTTGAAATA AACCTTCCTT TTTTATATGT TAATAAGGGG CCTTAGCTCA 240
 GCTGGGAGAG CGCCTGCTT GCACGCAGGA GGTCAAGGGT TCGATCCGC TAGGCTCCAC 300
 CAAAGATAGT TTGTTCTTG AAAACTAGAT AAGAAAAGTT AGTGTAAAAA GACGAAGAGA 360
 40 AACCGTAGGT TTTCTTCAA CCAAAACCGA GAAAGAATCT TTCCGTTTC ATAAGCGATC 420
 GCACGTTAT GAAAACACAA CAACACCTTC GTAAGAAGGA TGA 463

(2) INFORMATION FOR SEQ ID NO: 216:

- 45 (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 19 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

50 (ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

55 (iii) ANTI-SENSE: NO

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 216:

TGGCCGGTGC AAAGGGCTG

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Claims

- 15 1. Method for the detection and identification of at least one strain of Staphylococcus species or for the simultaneous detection of several microorganisms of which at least one strain of Staphylococcus species in a sample, comprising the steps of:

- 20 (i) releasing, isolating and/or concentrating the polynucleic acids from the micro-organism(s) to be detected in the sample;
 (ii) if need be amplifying the 16S-23S rRNA spacer region, or a part of it, from the micro-organism(s) to be detected, with at least one suitable primer pair;
 (iii) hybridizing the polynucleic acids of step (i) or (ii) to at least one of the following probes:

25 STAU-ICG 1 : TACCAAGCAAAACCGAGTGAATAAAGAGTT (SEQ ID NO 53)

STAU-ICG 2 : CAGAAGATGCGGAATAACGTGAC (SEQ ID NO 54)

STAU-ICG 3 : AACGAAGCCGTATGTGAGCATTGAC (SEQ ID NO 55)

30 STAU-ICG 4 : GAACGTAACCTCATGTTAACGTTGACTTAT (SEQ ID NO 56)

or to equivalents of said probes,
 and/or to any probe derived from SEQ ID NO 139, 140, 141, 142, 143 or 144 provided said probe hybridizes specifically to Staphylococcus species;
 35 (iv) detecting the hybrids formed in step (iii);
 (v) identification of the micro-organism(s) present in the sample from the differential hybridization signals obtained in step (iv).

- 40 2. Method according to claim 1 to detect and identify one or more Staphylococcus aureus strains, wherein step (iii) comprises hybridizing to at least one, and preferably both of the following probes:

45 STAU-ICG 3 : AACGAAGCCGTATGTGAGCATTGAC (SEQ ID NO 55)

STAU-ICG 4 : GAACGTAACCTCATGTTAACGTTGACTTAT (SEQ ID NO 56),

50 or to equivalents of said probes,
 and/or to any probe derived from SEQ ID NO 139, 140, 141, 142 or 143 provided said probe hybridizes specifically to Staphylococcus aureus.

- 55 3. Method according to claim 1 to detect and identify one or more Staphylococcus epidermidis strains in a sample, wherein step (iii) comprises hybridizing to any probe derived from SEQ ID NO 144 provided said probe hybridizes specifically to Staphylococcus epidermidis.

4. Method according to claim 1 wherein step (iii) is further characterized that the polynucleic acids of step (i) or (ii) are hybridized with a set of probes comprising at least two probes under the same hybridization and wash conditions, with said probes being selected from the sequences of table 1a or equivalents thereof, and/or from taxon-specific probes derived from any of the spacer sequences as represented in figures 1-103, with said taxon-specific

probe being selected such that it is capable of hybridizing under the same hybridization and wash conditions as at least one of the probes of table 1a.

5. Method according to claim 4, wherein the sample is originating from the respiratory tract and wherein step (iii) is further characterized that the set of probes comprises at least one probe chosen from the following spacer probes:

MYC-ICG-1 :	ACTGGATAGTGGTTGCGAGCATCTA	(SEQ ID NO 1)
MYC-ICG-22 :	CTTCTGAATAGTGGTTGCGAGCATCT	(SEQ ID NO 2)
MTB-ICG-1 :	GGGTGCATGACAACAAAGTTGGCCA	(SEQ ID NO 3)
MTB-ICG-2 :	GACTTGTCCAGGTGTTGTCCCAC	(SEQ ID NO 4)
MTB-ICG-3 :	CGGCTAGCGGTGGCGTGTCT	(SEQ ID NO 5)
MAI-ICG-1 :	CAACAGCAAATGATTGCCAGACACAC	(SEQ ID NO 6)
MIL-ICG-11 :	GAGGGGTTCCCGTCTGTAGTG	(SEQ ID NO 7)
MIL-ICG-22 :	TGAGGGGTTCTCGTCTGTAGTG	(SEQ ID NO 8)
MAC-ICG-1 :	CACTCGGTGATCCGTGTGGA	(SEQ ID NO 9)
MAV-ICG-1 :	TCGGTCCGTCCGTGTGGAGTC	(SEQ ID NO 10)
MAV-ICG-22 :	GTGGCCGGCGTTCATCGAAA	(SEQ ID NO 11)
MIN-ICG-1 :	GCATAGTCCTTAGGGCTGATGCGTT	(SEQ ID NO 12)
MIN-ICG-2 :	GCTGATGCGTTCGTCGAAATGTGTA	(SEQ ID NO 13)
MIN-ICG-22 :	CTGATGCGTTCGTCGAAATGTGT	(SEQ ID NO 14)
MIN-ICG-222 :	TGATGCGTTCGTCGAAATGTGT	(SEQ ID NO 15)
MIN-ICG-2222 :	GGCTGATGCGTTCGTCGAAATGTGTA	(SEQ ID NO 16)
MAL-ICG-1 :	ACTAGATGAACCGCGTAGTCCTTGT	(SEQ ID NO 17)
MHEF-ICG-1 :	TGGACGAAAACCGGGTGCACAA	(SEQ ID NO 18)
MAH-ICG-1 :	GTGTAATTCTTTAACTCTTGTGTAAAGTAAGTG	

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			(SEQ ID NO 19)
5	MCO-ICG-11 :	TGGCCGGCGTGTTCATCGAAA	(SEQ ID NO 20)
	MTH-ICG-11 :	GCACTTCAATTGGTGAAGTGCGAGCC	(SEQ ID NO 21)
	MTH-ICG-2 :	GCGTGGTCTTCATGCCCGG	(SEQ ID NO 22)
10	MEF-ICG-11 :	ACCGTGGTCCTCGTGG	(SEQ ID NO 23)
	MSC-ICG-1 :	TCGGCTCGTCTGAGTGGTGTC	(SEQ ID NO 24)
	MKA-ICG-1 :	GATGCGTTGCTACGGTAGCGT	(SEQ ID NO 25)
	MKA-ICG-2 :	GATGCGTTGCCCTACGGTAGCGT	(SEQ ID NO 26)
15	MKA-ICG-3 :	ATGCGTTGCCCTACGGTAGCGT	(SEQ ID NO 27)
	MKA-ICG-4 :	CGGGCTCTGTCGAGAGTTGTC	(SEQ ID NO 28)
	MKA-ICG-5 :	CCCTCAGGGATTTCTGGGTGTTG	(SEQ ID NO 182)
20	MKA-ICG-6 :	GGACTCGTCCAAGAGTGTGTC	(SEQ ID NO 183)
	MKA-ICG-7 :	TCGGGCTTGGCCAGAGCTGTT	(SEQ ID NO 184)
	MKA-ICG-8 :	GGGTGCGAACAGCAAGCGA	(SEQ ID NO 185)
25	MKA-ICG-9 :	GATGCGTTGCCCTACGGG	(SEQ ID NO 186)
	MKA-ICG-10 :	CCCTACGGGTAGCGTGTCTTTG	(SEQ ID NO 187)
	MCH-ICG-1 :	GGTGTGGACTTGACTTCTGAATAG	(SEQ ID NO 29)
30	MCH-ICG-2 :	CGGCAAAACGTCGGACTGTCA	(SEQ ID NO 30)
	MCH-ICG-3 :	GGTGTGGCCTTGACTTATGGATAG	(SEQ ID NO 210)
	MGO-ICG-1 :	AACACCCTCGGGTGCTGTCC	(SEQ ID NO 31)
35	MGO-ICG-2 :	GTATGCGTTGTCGTCGCGGC	(SEQ ID NO 32)
	MGO-ICG-5 :	CGTGAGGGTCATCGTCTGTAG	(SEQ ID NO 33)
	MUL-ICG-1 :	GGTTTCGGGATGTTGCCACC	(SEQ ID NO 175)
40	MGV-ICG-1 :	CGACTGAGGTCGACGTGGTGT	(SEQ ID NO 176)
	MGV-ICG-2 :	GGTGTGAGCATTGAATAGTGGTTGC	(SEQ ID NO 177)
	MGV-ICG-3 :	TCGGGCCGCGTGTCTCACCAAA	(SEQ ID NO 211)
45	MXE-ICG-1 :	GTTGGGCAGCAGGCAGTAACC	(SEQ ID NO 178)
	MSI-ICG-1 :	CCGGCAACGGTTACGTGTT	(SEQ ID NO 179)
	MFO-ICG-1 :	TCGTTGGATGGCCTCGCACCT	(SEQ ID NO 180)
50	MFO-ICG-2 :	ACTTGGCGTGGGATGCCGGAA	(SEQ ID NO 181)
	MML-ICG-1 :	CGGATCGATTGAGTGCTTGTCCC	(SEQ ID NO 188)
	MML-ICG-2 :	TCTAAATGAACGCACTGCCGATGG	(SEQ ID NO 189)
55	MCE-ICG-1 :	TGAGGGAGCCCGTGCCTGTA	(SEQ ID NO 190)

MHP-ICG-1 :	CATGTTGGGCTTGATCGGGTGC	(SEQ ID NO 191)
PA-ICG 1 :	TGGTGTGCTGCGTGATCCGAT	(SEQ ID NO 34)
5 PA-ICG 2 :	TGAATGTTCGTGGATGAACATTGATT	(SEQ ID NO 35)
PA-ICG 3 :	CACTGGTGATCATTCAAGTCAG	(SEQ ID NO 36)
PA-ICG 4 :	TGAATGTTCGT(G/A)(G/A)ATGAACATTGATTCTGGTC	
10		(SEQ ID NO 37)
PA-ICG 5 :	CTCTTCACTGGTGATCATTCAAGTCAG	(SEQ ID NO 38)
MPN-ICG 1 :	ATCGGTGGTAAATTAAACCCAAATCCCTGT	(SEQ ID NO 49)
15 MPN-ICG 2 :	CAGTTCTGAAAGAACATTCCGCTTCTTC	(SEQ ID NO 50)
MGE-ICG 1 :	CACCCATTAATTTCGGTGTAAAACCC	(SEQ ID NO 51)
20 Mycoplasma-ICG :	CAAAACTGAAAACGACAATCTTCTAGTTCC	(SEQ ID NO 52)
STAU-ICG 1 :	TACCAAGCAAAACCGAGTGAATAAAGAGTT	(SEQ ID NO 53)
STAU-ICG 2 :	CAGAAGATGCGGAATAACGTGAC	(SEQ ID NO 54)
25 STAU-ICG 3 :	AACGAAGCCGTATGTGAGCATTTGAC	(SEQ ID NO 55)
STAU-ICG 4 :	GAACGTAACCTCATGTTAACGTTGACTTAT	(SEQ ID NO 56)
ACI-ICG 1 :	GCTTAAGTGCACAGTGCTCTAAACTGA	(SEQ ID NO 57)
30 ACI-ICG 2 :	CACGGTAATTAGTGTGATCTGACGAAG	(SEQ ID NO 58)

or equivalents of said probes,

and/or wherein the set of probes comprises at least one taxon-specific probe derived from the spacer region sequence corresponding to one of the micro-organisms to be detected in said sample, said spacer region sequence being chosen from any of the sequences as represented by SEQ ID NO 76 to 106, 157 to 174, 124, 125, 111 to 115, 139 to 144, or 126 to 130,

and with said probes or equivalents being possibly used in combination with any probe detecting at least one of the following organisms: Haemophilus influenzae, Streptococcus pneumoniae, Moraxella catarrhalis or Bordetella pertussis.

- 40 6. Method according to claim 4, wherein the sample is originating from food, and wherein the set of probes as defined in step (iii) comprises at least one probe chosen from the following spacer probes:

LIS-ICG 1 :	CAAGTAACCGAGAATCATCTGAAAGTGAATC	(SEQ ID NO 39)
LMO-ICG 1 :	AAACAAACCTTACTTCGTAGAAGTAAATTGGTTAAG	
50 LMO-ICG 2 :	TGAGAGGTTAGTACTTCTCAGTATGTTGTT	(SEQ ID NO 41)

LMO-ICG 3 :	AGGCACTATGCTGAAGCATCGC	(SEQ ID NO 42)
LIV-ICG 1 :	GTTAGCATAAATAGGTAACATTATGACACAAGTAAC	
5		(SEQ ID NO 43)
LSE-ICG 1 :	AGTTAGCATAAGTAGTGTAACTATTATGACACAAG	(SEQ ID NO 44)
10		
LISP-ICG 1:	CGTTTCATAAGCGATCGCACGTT	(SEQ ID NO 212)
STAU-ICG 1 :	TACCAAGCAAAACCGAGTGAATAAAGAGTT	(SEQ ID NO 53)
15		
STAU-ICG 2 :	CAGAAGATGCGGAATAACGTGAC	(SEQ ID NO 54)
STAU-ICG 3 :	AACGAAGCCGTATGTGAGCATTGAC	(SEQ ID NO 55)
20		
STAU-ICG 4 :	GAACGTAACCTCATGTTAACGTTGACTTAT	(SEQ ID NO 56)
BRU-ICG 1 :	CGTGCCGCCCTCGTTCTCTTT	(SEQ ID NO 59)
25		
BRU-ICG 2 :	TTCGCTTCGGGTGGATCTGTG	(SEQ ID NO 60)
BRU-ICG 3 :	GCGTAGTAGCGTTGCGTCGG	(SEQ ID NO 193)
30		
BRU-ICG 4 :	CGCAAGAAGCTTGCTCAAGCC	(SEQ ID NO 194)
SALM-ICG 1 :	CAAAACTGACTTACGAGTCACGTTGAG	(SEQ ID NO 61)
35		
SALM-ICG 2 :	GATGTATGCTTCGTTATTCCACGCC	(SEQ ID NO 62)
STY-ICG 1 :	GGTCAAACCTCCAGGGACGCC	(SEQ ID NO 63)
40		
SED-ICG 1 :	GCGGTAATGTGTGAAAGCGTTGCC	(SEQ ID NO 64)
YEC-ICG 1 :	GGAAAAGGTACTGCACGTGACTG	(SEQ ID NO 198)
45		
YEC-ICG 2 :	GACAGCTGAAACTTATCCCTCCG	(SEQ ID NO 199)
YEC-ICG 3 :	GCTACCTGTTGATGTAATGAGTCAC	(SEQ ID NO 200)

or equivalents of said probes,

and/or wherein the set of probes comprises at least one taxon-specific probe derived from the spacer region sequence corresponding to one of the micro-organisms to be detected in said sample, said spacer region sequence being chosen from any of the sequences as represented by SEQ ID NO 116, 117, 118-121, 213-215, 139-144, 131, 132, 154, 133-138, 195 or 196,

with said probes or equivalents being possibly used in combination with any probe detecting strains of Campylobacter species.

- 45
7. Composition comprising at least one of the probes as defined in claims 1 to 3.
 8. Probe as defined in any of claims 1 to 3.
 - 50 9. Reverse hybridization method according to any of claims 1 to 6 wherein the probes are immobilized on a known location on a solid support, more preferably on a membrane strip.
 10. Kit for the detection and identification of at least one strain of *Staphylococcus* species, or the simultaneous detection and identification of several micro-organisms of which at least one strain of *Staphylococcus* species in a sample, comprising the following components:
 - 55 (i) when appropriate, at least one suitable primer pair to allow amplification of the 16S-23S rRNA spacer region, or a part of it;

- (ii) at least one of the probes as defined in claim 8;
(iii) a buffer, or components necessary to produce the buffer, enabling a hybridization reaction between said probes and the polynucleic acids present in the sample, or the amplified products thereof;
5 (iv) a solution, or components necessary for producing the solution, enabling washing of the hybrids formed under the appropriate wash conditions;
(v) when appropriate, a means for detecting the hybrids resulting from the preceding hybridization.

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Figure 1

AAGGAGCACC ACGAAAACGC CCCAACTGGT GGGGCCGTAGG CCGTGAGGGG TTCTTGTCTG TAGTGGGGGA
GAGCCGGGTG CATGACAACA AAGTTGGCCA CCAACACACT GTTGGGTCTT GAGGCAACAC TCGGAACTTGT
TCCAGGTGTT GTCCCCACCGC CTTGGTGGTG GGGTGTGGTG TTTGAGAACT GGATACTGGT TGGCAGGCATC
AATGGATAACG CTGCCGGCTA GCGGTGGGT GTTCTTTGTG CAATATTCTT TGTTTTTGT TGTGT

(SEQ ID NO 76)

Figure 2

AAGGAGGCC ACCAAAAGCA CCCCAACTGG TGGGGTGGCA GCCGTGAGGG GTTCGGGTCT GTAGTGGACG
GGGGCCGGNT GCGCAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAACAC TCGGTCCGTC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG GTGTGGTGT TGAGTATTGG ATAGTGGTTG CGAGCATCTA
GATGAGGCA TGGTCTTCGT GGCGGGCGTT CATCGAAATG TGTAAATTCT TCCTTAACTC TTGTGTGT

(SEQ ID NO 77)

Figure 3

AAGGAGGCACC ACGAAAGCA CCCCAAACCTGG TGGGGTGGGA GCCGTGAGGG GTTCCCGTCT GTAGTGGACG
GGGGCCGGGT GCGCAACAGC AAATGATTGC CAGACACACT ATTGGCCCT GAGACAAACAC TCGGGTCCGTC
CGTGTGGAGT CCCTCCATCT TGTTGGTGG GTGTGGTGT TGAGTATTGG ATAGTGGT'TG CGAGCATCTA
GATGAGGCCA TGGTCTTCGT GGCGGGCGTT CATCGAAATG TGTAAATTCT TTGTAACCTC TTGTTAACTC TTGTTGTGT

(SEQ ID NO 78)

Figure 4

AAGGAGCACC ACGAAAAGCA CTCCAAATTGG TGGGTGGGA GCCGGTGGCT GTAGTGGACG
GGGGCCGGNT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAAACAC TCGGTGATC
CGTGTGGAGT CCCTCCCATCT TGTTGGTGG GTGTGGTGT TGAGTATTGG ATAGTGGTGG CGAGGCATCTA
GATGAGGCA TAGTCCTTGT GGCTGTGATGCG CTCGTCGAAA TGTGTAATT CTTCTTTGGT GTNTGGTGT

(SEQ ID NO 79)

Figure 5

AAGGAGGCC ACC GCGAAAAGCA TCCC CAATTGG TGGGGTGGGA GCCGTGAGGG GTTAGTGGTCT GTAGTGGACG
AAAACCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAACAC TCGGTGATC
CGTGTTGAGT CCCTTCATCT TGTTGGTGG TGAGTATTGG ATAGTGGTTC CGAGGCATCTA
GATGAGGGCG TAGTCCTTTG TGGCTGATGC GTTCATCAAAT GTGTAAATT TCTTTGG TTNTNTGTGTG
T

(SEQ ID NO 80)

Figure 6

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGGGA GCCGTGAGGG GTTCGGGTCT GTAGTGGACG
GGGGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAAACAC TCGGTGATC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG GTGTGGTGT TGAGTATTGG ATAGTGGTT CGAGCATCTA
GATGAGGCA TAGCCCTTGC GGCTGATGGC TTCGNCGAA TGTGTAATT TTCTCTGGT TTCTGTGT

(SEQ ID NO 81)

Figure 7

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCAGA GCGGTGAGGG GTTCTCGTCT GTAGTGGACG
GNAGCCGGGT GCACAAACAGC AAATGATTCG CAGACACACT ATTGGGCCCT GAGAACAC TCGGTCGATC
CGTCTGGAGT CCCTCCATCT TGGTGGTGG TGAGTATTGG ATAGTGGTTG CGAGCATCTA
GATGAGCGCG TAGTCCTTCG TGGCTGATGC GTTCATCGAA ATGTGTAATT TCTTCTTTGG TTTTGGGTGT
GT

(SEQ ID NO 82)

Figure 8

AAGGAGGCC ACCGAAAAGCA CTCCCAATTGG TGCGGTGGGA GCCGGTGAAGGG GTAGCTGGTCT GTAGCTGGACG
GGGGCCGGGT GCACAACAGC AAATGATCGC CAGACACACT ATTGGCCCT GAGACAACAC TCGGTGATCG
CGTGTGGAGT CCCTCCATCT TGGTGGTGG TGACTATTGG ATAGTGGTT CGAGGCATCTA
GATGAGGCCA TAGTCCTTGT GGGCTGATGT GTTTCATCAA AATGTTAAAT TTCTTTNGTGT GTTNTNGTGT
GT

(SEQ ID NO 83)

Figure 9

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTAGTGGACG
GGAGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAAACAC TCGGGTCGATC
CGTGGGAGT CCCTCCATCT TGGTGGTGG TGTTGGTGT TGAGTATTGG ATAGTGGTGG CGAGGCATCTA
GATGAGGGCG TAGTCCTTCG TGGCTGATGC GTTCAATTGAA ATGTCATTGAA TCTTCCTCTGG TTTTTGTGTG
T

(SEQ ID NO 84)

Figure 10

AAGGAGGCC ACCAAAAGCA CTCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTTCCCGTCT GTAGTGGACG
GGGGCCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGCCCT GAGACAAAC CTCGGTCCGATC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG TGAGTATTGG ATAGTGGTTC CGAGCACTA
GATGAGGCA TAGTCCTTGT GGCTGATGCG CTCGTGAAA TGTGTAATT CTTCTTTGGT TTTTGTGTGT

(SEQ ID NO 85)

Figure 11

AAGGGAGCACC ACGAAAAGCA CTCCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTAGTCGTCT GTTCCCGTCT
GGGGCCGGGT GCGAACAGC AAATGATTGC CAGACACACT ATTGGCCCT GAGACAACAC TCGGTGATC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG GTGTTGGTGT TTGAGTATTG GATAAGGGT GCGAGCATCT
AGATGAGGCC GTAGTCCTTG TGGCTGATGC GTTCGTCGAA ATGTTGAATT TCTTCCTTGG GTTTTGTGT
GT

(SEQ ID NO 86)

Figure 12

AAGGAGCACC ACGAAAAGCA CCCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTTCTCGTCT GTAGTTGGACCG
GNAGCCGGNT GCGCZAACAGC AAATGATTCG CAGACACACT ATTGGGCCCT GAGACAACAC TCGGNCGATC
CGTGTGGAGT CCCTCCATCT TGCTGGTGG GTGTNGTGT TGAGTATGG ATAGTGGTTG CGAGCATCTA
GATGGGGCGCG TAGTCCCTTG TGACTGATGC GTTCATCAA ATGTGTAATT TCTTTTTGN NTITTINGTGTG
T

(SEQ ID NO 87)

Figure 13

AAGGAGGCC ACAGAAAAGCA CTCCAATTGG TGGGGTGGGA GCCGTGAGGG GTAGTGGACG
GGAAACCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGCCCT GAGACAACAC TCGGTGGATC
CGTGTGGAGT CCCTTCCATCT TGGTGGTGGG GTGTGGTGT TGAGTATTGG ATAGTGGGTG CGAGCATCTA
GATGAGCGCA TAGTCCTTG TGGCTGACGC GTTCATCGAA ATGTGTAATT TCTTCTTTGG TTTTGTGTG
T
(SEQ ID NO 88)

Figure 14

AAGGAGGCACC ACCGAAAGCA CTCCCAATTGG TGGGGTGGGA GCCGTGANGG GTTCCCGGTCT GTAGTGGACCG
GGGGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAACAC TCGGTGCGATC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG TGAGTATTGG ATAGTGGTTC CGAGCATCTA
GATGAGGCA TAGTCCTTAG GGCTGTGCG TTCGTGATGCG TTGTTAATT TTCTCTTGTT TTTTGTGTGT

(SEQ ID NO 89)

Figure 15

AAGGAGGCC ACCGAAAAGCA TCCCAATTGG TGGGTGGCGA GCGGTGAGGG GTTCTCGTCT GTAGTGGACG
AAACCCGGGT GCACAAACAGC AAATAATTGC CAGACACACT ATTGGGCCCT GAGACAAACAC TCGGTGATC
CGTGTGGGT CCCTTCCATCT TGGTGGTGGG GTGTGGTGT TGAGTATGG ATAGTGGTTG CGAGCATCTA
GATGAACGG TAGTCCTTCG TGGCTGACGT GTTCATCGAA ATGTGTAATT TCTTNNTTA ACTCTTGTGT
GT

(SEQ ID NO 90)

Figure 16

AAGGAGCACC ACGAAAAGCA CCCCAATTGG TGGGGTGCAGA GCCGGTGAGGG GTTCTCGTCT GTAGTGGACG
GGAGCCGGGT GCACAACAGC AAATGATGCC CAGACACACT ATTGGGCCCT GAGACAAACAC TCGGTCAAGTC
CGTGTGGTGT CCCTCCATCT TGGTGGTGGG GTGGGGTGT TGAGTATTGG ATAGTGGTTG CGAGCATCTA
GATGAACGGG TAGTCCTTGT GACTGACGTTG TTCATCGAAA TGTGTAATTI CTTTCTAAC TCTTGTGT

(SEQ ID NO 91)

Figure 17

AAGGAGGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGGCA GCCGTGAGG GTTAGTCGTCT GTAGTGGACG
AAAGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGCCCT GAGACAAACAC TCGGGTGAAC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG TGAGTGTGTT TGAGTATTGG ATAGTGGTGT CGAGCATCTA
GATGAAACGCG TGGTCTTCAT GGCGGGGTG TTCAATCGAAA TGTGTAATA TGTGTAATAT CTTCTCTGGT TTTCGGTGTG
T

(SEQ ID NO 92)

Figure 18

AAGGAGGCC ACCGAAAGCA CTTCAATGG TGAAGTGGCA GCCGTGAGGG GTAGTGGACG
AAAACCGGNT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGCCCT GAGACAAACAC TCGGTGATC
CGTGTGGAGT CCCTCCATCT TGGTGGGG TGCTGGTGT TGAGTATTGG ATAGTGGTGG CGAGCAATCTA
GATGAAACGGG TGGTCTTCAT GGCCGGCGTG TTCATCGAAA TGTGTAATT CTTTTNNAC TCTTGTGT

(SEQ ID NO 93)

Figure 19

AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCGGTGAGGG GTTCTCGTCT GTAGTGGACG
AAAGCCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAACAC TCGGTGCGAAC
CGTGTGGAGT CCCTCCATCT TGGTGGTGGG GTGTGGTGT TGAGTATTGG ATAGTGGTGG CGAGCATCTA
GATGAAACGGCG TGGTCTTCAT GGCGGGGTG TTCATCGAAA TGTGTAATT CTTCTTGGT TTNGTGTGT

(SEQ ID NO 94)

Figure 20

AAGGAGGCC ACCAAAAAGCA CTTCAATTGG TGAAGTGGCGA GCGGTGAGGG GTTCTCGTCT GTAGTGGACG
AAAACCGGT GCACAACAGC AAATGATTGC CAGACACACT ATGGGCCCT GAGACAACAC TCGGTGCGATC
CGTGTGGAGT CCCTCATCT TGGTGGTGG TGAGTATTGG ATAGTGGTTG CGAGCCATCTA
GATGAACGCG TAGTCCTTCG NGGNNCNGCGT GTTCATCGAA ATGTGTAATT TCTNTNTAA CTCTNGTGTG

T

(SEQ ID NO 95)

Figure 21

AAGGAGGCC ACCAAAAGCA TCCCATTGG TGGGGTGTGA GCCGGTGAGG GTAGCTCGTCT
AAAACCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGGCCCT GAGACAACAC TCGGGTCGATC
CGTGTGGAGT CCCTTCCATCT TGTTGGTGG TGAGTATTGG ATAGGGTTG CGAGCATCTA
GATGAAACGGG TAGTCCCTTCG GGGCCGGCGT GTTCAATCGAA ATGTTAAATT TCTTTTTAA CTCTTGTGTG
T

(SEQ ID NO 96)

Figure 22

AAGGAGCACC ACCAAAAAGCA CTTCANTTGG TGAAGTGGCA GCCGTGAGGG GTTCTCGTCT GTAGTGGACG
AAAACCGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGCCCT GAGACAACAC TCGGTGAAAC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG TGAGTATTGG ATAGTGGTTG CGAGCATCTA
GATGAAACGGG TGGTCTTCAT GGGCGGGTG TTCATCGAAA TGTGTAATT CTTCTTTAAC TCTTGTGTGT

(SEQ ID NO 97)

Figure 23

AAGGAGCACC ACGAAAAGCA CTTCAATTGG TGAAGTGCAGA GCCGTGAGGG GTTCTCGTCT GTAGTGGACG
AAAACCGGGT GCACAAACAGN AAATGATTGC CAGACACACT ATTGGGCCCT GAGACAACAC TCGGGTCGATC
CGTGTGGAGT CCCTCCATCT TGCTGGTGG GTGTGGTGT TGAGTATTGG ATAGTGGTGG CGAGCATTCA
GATGAAACGGCG TGGTCTTCAT GGCCNGCGTG TTCATCGAAA TGTGTAATT CTTTTAAC TCTTGTGT

(SEQ ID NO 98)

Figure 24

AAGGAGGCC ACCGAAAGCA CTTCAATTGG TGAAGTGGCA GCCGTGAGGG GTTCTCGTCT GTAGTGGACG
AAAACGGGT GCACAAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGACACAC TCGGTGGATC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG TGAGTATTGG ATAGTGGTGG CGAGCATCTA
GATGAAACGCG TGGTCTTCAT GGCCGGCGTG TTCATCGAAA TGTGTAATT CTTTTTAAC TCTTGTGT

(SEQ ID NO 99)

Figure 25

AAGGAGCACC ACGAAAAGCA CCCCAACTTGG TGGGGTGGGA
GGGGCCGGGT GCACAACAGC AAATGATTGC CAGACACACT ATTGGGCCCT GAGGCAACAC TCGGCTCGTT
CTGAGTGGTG TCCCCTCCATC TTGGTGGTGG GGTGTTGGTGT TTGAGTATG GATAAGTGGTT GCGAGGCAATCT
AAACGGATGC GTGGCCGGCA ACGGGTGGCGT GTTCGTTGAA ATGTGTAAATT TCTTTTGTGG TTTTTGTGTG
T

(SEQ ID NO 100)

Figure 26

AAGGAGGCC ACCGAAAGCA TCCCAACAAG TGGGGTGC_{AA} NCCGTGAGGG GTAGTGGACG
AAAGCCGGGT GCACGACAAC AAGCAAAGCC AGACACACTA TTGGGTCTG AGGCAACACT CGGGCTCTGT
TCGAGAGTTG TCCCACCATC TTGGTGGTGG GGTGTGGTGT TTGAGAATTG GATAAGTGGTT GCGAGGCATCA
AATGGATGCG TTGCCCTACG GGTAGCGTGT TCTTTGTGC AATTATTC TTGGTTTTT GTGT

(SEQ ID NO 101)

Figure 27

AAGGAGCACC ATTTCCCACT CGATGAACTA GGGAACATAA AGTAGGCATC TGTAGTGGAT ATCTACTTGG
TGAATATGTT TTGTAAATCC TGTCACCCCC GTGGATGGGT AGTGGCAAAC ACGTGGGACT GTCATAGAA
TTGAAACGCT GGCACACTGT TGGGTCTCTGA GCGAACACGT TGTGTGTGCA CCCTGCTTGG TGGTGGGGTG
TGGACTTTGA CTTCTGAATA GTGGTTGCCA GCATCTAAC ATAGCCTCGC TCGTTTCGA GTGGGGCTGG
TTTGCAATT TTA

(SEQ ID NO 102)

Figure 28

AAGGAGGCC ACC ATTTC CCA GTT CGG ATG TGA ACT AGGGAA CATA AAG TAG GGC AT CTGTAG TGGG TATCTACTTG
GTGAATATGT TTT GTAAATC CTGTCCACCC CCGTGGATGG GTAGTCGGCA AACACGTGGCA AAAGCTGGGA CTGTCTAAAG
AATTGAAACG CTGGCACACT GTTGGGT CCT GAGGCAACAC GTTGTGTGT CACCCCTGCTT GGTGGTGGGG
TGTGGACTTT GACTTCTGAA TAGTGGTTGC GAGGCATCTAA ACATAGCCTC GCTCGTTTC GAGTGAGGCT
GGTTTTGGCA ATTTTA

(SEQ ID NO 103)

Figure 29

AAGGAGCACC ACGAAGAGCA CTCCAATTGG TGGGGTGGGA GCGGTGAGGG GTCATCGTCT GTAGTGACG
AAGACCGGGT GCACGCCAAC AAGCTAACCC AGACACACTA TTGGGTCCCTG AGGCAACACC CTCGGGGTGGCT
GTCCCCCAT CTTGGTGGTG GGGTGTGGTG TTTGAGAATT GGATAGTGGT TGGCAGACATC AAAATGTATG
CGTTGTCGTT CTGGCAACCG TGTTCTTTT GTGCAATTAA TTCTTTGGTT TTTGTAGTGT TTGT

(SEQ ID NO 104)

Figure 30

AAGGAGCACC ACGAAGGAGCA CTCCAATTGG TGGGGTGCAGA GCGCNGAGGG GTCATCGTCT GTAGTGGACG
AAGACTGGGT GCACGACAAC AAAGCAAGCC AGACACACTA TTGGGTCCCTG AGGCAACACCC CTCGGGGTGGCT
GCCCTCCAT CTTGGTGGTG GGGTGTGGTG TTTGAGAACT GGATAGTGGT TGGGAGGCATC AAAAATGTAT
GCGTGTGCGT TCGGGACAAC GTGTTCTTT TGTGCAATT TAATTCTTT GGTTTGGTA GTGTTGT

(SEQ ID NO 105)

Figure 31

AAGGAGCACC ACGAGGAAGCA CTCCAATTGG TGGGGTGC_{AA} GCCGGTGAGGG GTCATCGTCT GTAGTGACG
AAGACCCGGGT GCACGACAAAC AAGCAAAGCC AGACACACTA TTGGGTCC_{CTG} AGGCAACACC CTCGGGTGCT
GTCCCCCAT CTTGGTGGTG GGGTAGGAACT TTTGAGGTGGT GGATAGTGGT TGGGAGC_{ATC} AAATGTATG
CGTTGTGTT CGGGCAACG TGTTCTTTT GTGCCAATT TATTCTTGG TTTTTAGT GTTTGT

(SEQ ID NO 106)

Figure 32

AAGGAGGCC ACCGAAAGCA CCCCAATTGG TGGGGTGCCTA GCCGTGAGGG GTTCCCGCCT GTAGTGGGG
GGGCCGGGTG CGCAACAGCA AATGATTGCC AGACACACTA TGGGCCCTG AGGCAACACT CGGATCGATT
GAGTGCTTGT CCCCCCATCT TGGTGTGGG GTGTGTGTT TGAGAACTGG ATAGTGGTG CGAGCATCTA
AATGAACGCA CTGCCGATGG TGTTGTGTA GTTTTGTGTA ATTATTATTCT TTGGTTTTG TGTTGT

(SEQ ID NO 107)

Figure 33

AAGGAGGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGGCGA GCCGTNAGGG GTTCTCGTCT GTAGTGGATG
GCAGCCGGGT GCACANCAGC AAATGATTGC CAGACZACACT ATTGGGCCCT GAGACAACAC TCGGTCAAGTC
CGTGTGGAGT CCCTCCATCT TGGTGGTGG TGTTGNGTT TGAGTATTGG ATAGTGGTTG CGANCATCTA
GATGAACGGC TAGTCTCNG TGGCTGACGT GTTCAATCAA ATGTGTAATT TCTTTTANGG GTTTNGGTGT
CT

(SEQ ID NO 108)

Figure 34

AAGGAGGCC ACCGAAAGCA CTCCAATTGG TGGGGTGCAGA GCGGNGAGGG GTTCTCGCCT GTAGTGGNCG
AGGGCCGGAT GCACAAAC ACATGATTGC CAGACACACT ATTGGCCCT GANACAAACAC TCGGCCAGTC
CGTGTGGTGT CCCTCCATCT TGTTGGTGGG GTGTGGTGT TGAGTATNGG ATAGTNGTGA NGANCATCTA
AACGGCTGCG TNGNCNNGAA CGGTGGCGTG TTCGNTAAA TGTGTAATT CTTTTNNGGT TTGGGTGTNT

(SEQ ID NO 109)

Figure 35

AAGGAGCACC ACGAAAAGCA CTCCAATTGG TGGGGTGCAGA GCCGTGAGGG GTTCTGGCCT GTAGTGGCG
ANGGCCGGGT GCACAAACAA ACATGATTGC CAGACACACT ATTGGCCCT GAGACAACAC TCGGCCAGTC
CGTGTTGGTCCCNCATCT TGTTGGTGGG GTGTGGTGT TGAGTATTGG ATAGTGGTGC CGAGGCATCTA
AANGGNTGGCTTGCCGNAN CNGTGGCGTN TTCGNTAAA TGTGTAANTT CTTTTTNGGT TTGTGTGTGT
(SEQ ID NO 110)

Figure 36

ATCGAAGATC CCGGCTTCTT CATAAGCTCC CACACCAATT GCTTGATTCA CTTGGTAGAC GATTGGGTCT
GTAGCTCACT TGGTTAGACC GCACCCCTGA TAAGGGTGA GTCGGGCAGTT CGAATCTGCC CAGACCCACC
AATTGTTGGT GTGCTGGTG ATCCGATACT GGGCCATAGC TCAGGCTGGGA GAGGGCCTGC TTGCAACGCA
GGAGGTCAAG AGTTCACTCC TCCTTGGCTC CACCATCTAA AACAACTCGTC GAAAGGCTCAG AAATGAAATGT
TCGGTGGATGA ACATTGATT CTGGTCTTTG CACCAACT GTTCTTTAAA AATTGGGTAA TGTGATAGAA
GTAAGACTGA ATGATCTCTT TCACGGTGA TCATTCAAGT CAAGGTAAA TTGGCAGTT CAAGGGCGAA
TTTTCGGGA ATGTCGTCTT CACAGTATAA CCAGGTTGCT TGGGGTTATA T

(SEQ ID NO 111)

Figure 37

ATCGAAGACA TCAGCTTCTT CATAAGTATC CACACGAATT GCTTGATTCA TAGTCGAACG AATGCTGTAA
CCGGACCCGT GTTATAGGTC TGTAGCTTCAG TTGGTTAGAG CGCACCCCTG ATAAGGGTGA GGTCTGGCAGT
TCAAATCTGC CCAGACCTAC CAATTGCTTG GTCGAGAAAGA ATACGGGCC ATAGCTCAGC TGGCAGAGCG
CCTGCCCTGCA ACGCAGGAGG TCAGCGGTC GATCCCGCTT GGCTCCACCA CTCTCTCGTG TTGGGGTGAG
TGTAAAGAG TTCAGAAATG ATGCCGCTTC AGGTTGTCC TGTTGACTGC TGATTCTGG TCTTTTGACC
GGTACGAAA TCGTTCTTA AAAATTGGA TATGTGATAG AAGTGAETGA TTAATTGCTT TCACTGGCAA
TTGATCTGGT CAAGGTTAAA TTGGTAGTTC TCAAGACGCC ATTTCGGCA GAATGTCGGC TTCACGGATTG
AGACAGAAC CAGATTGCTT GGGTTATAT

(SEQ ID NO 112)

Figure 38

ATCGAAGACA CCGGCTCTGT CATAAGCTCC CACACGAAATT GCTTGATTCA CTTGCGAAAG GCGATTGGT
TAGACCCGA GAGTAACGGAT TGGGTCTGTA GCTCAGTTGG TTAGAGGCC CCCCTGATAA GGGTGAAGGTC
GGCAGTCGA ATCTGCCAG ACCCACCAAT CGAAGGGGCC ATAGCTCAGC TGGGAGAGCG CCTGCTTTGC
ACGGAGGAGG TCAGCGGTT GATCCCCGCTT GGCTCCACCA TAACTCTAG TGCCCGAAAG CTCAGAAATG
AGTGTITACC AGGATGAGGT TGATTGCCCTG GGTGAAACAT TGATTTCTGG ACTTTGGCC AGAACTGTT
TTAAAAATT TGGGTATGTG ATAGAAGTAG ACCGATGTGT TGCTTTCACT GGCAGCATGT CGCGTCAAAGG
TAAATTTGC GTGTTCTCTA TGCAAATTG CGGCGAATTT CGTCTCAG TTATAGACAG TAACCAGATT
GCTTGGGGTT ATAT

(SEQ ID NO 113)

Figure 39

ATCGAAGACT TCAGCTTCTT CATAAGTTCC CACACGAATT GCTTGATTCA CTTGCGAAAA GCGGATTGGGT
TGAGACCCGA GAGTGACCGAT TGGGTCTGTA GCTCAGTGG TAGAGGGCA CCCCTGATAA GGGTGAGGTC
GGCAGTTCGA ATCTGCCAG ACCCACCAAAT TGTCGGGATG GCCAGTGTCA AATGGGGCCA TAGCTCAGCT
GGGAGGGCC CTGCTTTGCA CGCAGGGGT CAGGAGTTCG ATCCTCCTTG GCTCCACCAT CAACTCACGA
TCGCTGAAAG CTCAGAAATG AACATGGTA GTTCAATGTT GATTTCCTGGT CTTTGGCCA GAACTGTTCT
TTAAAAATT GGTTATGTGA TAGAAGTGGAC TAACAGGGTG TTTCACTGCA CGTTGTTAAT CAAGGCAAAA
TTTGCAGTCAAGCCGAA TTTCGGGA ATGTCCGTCTT CACGGTACGA ATCTATAACC AGATTGCTTG
GGTTATAT

(SEQ ID NO 114)

Figure 40

ATCGACGACA TCAGCTGTCT CATAAGCTCC CACACGAAATT GCTTGATTCA TTGAAGAAGA CGATTAGTT
AGCAACCTTC GATTGGTCT GTAGCTCACT TGTTAGAGC GCACCCCTGA TAAGGGTGTAG GTCGGCAGTT
CGAATCTGCC CAGACCACC AATTGCTGG GGCCATAGCT CAGCTGGGAG AGGCCCTGCC TTGCACGCC
GAGGTCAAGCG GTTCGATCCC GCTTGGCTCC ACCACCCCGC TTGCCAGCTT GTCAAAGCTT AGAAATGAAT
ATTCGCGTCG AATATTGATT TCTGAACCTT ATCAGAATCG TTCTTTAAAA ATTGGGTAT GTGATAAGAAA
GATAGACTGG ACAGCACCTT CACTGGTGTG TGTTCAAGGCT AAGGTAATAAT TTGTGAGTAA TTACAAGTT
TCGGCGAATG TTGTCTCAC AGTATAACCA GATTGCTGG CGTTATAT

(SEQ ID NO 115)

Figure 41

TAAGGAAAAG GAAACCTGTG AGTTTCGTT CTTCTCTGTT TGTCAGTT TGAGAGGTT ATTCTTCTCT
ATACTGTTG TTCTTTGAAA ACTAGATAAG AAAGTTAGTA AAGTTAGCAT AAATAGGTTAA CTATTATGA
CACAAGTAAC CGAGAAATCAT CTGAAAGTGA ATCTTTCATC TGATTGGAAAG TATCATCGCT GATAACGAAAAA
ATCAGAAAAA CAACCTTAC TTCATCGAAG TAAATT

(SEQ ID NO 116)

Figure 42

CTAAGCAAA CGAACCTGT GAGTTTCGT TCTTCTAT TTGTTTAGTT AGTACTTCTC
AGTATGTTG TTCTTGAAA ACTAGATAAG AAAGTTAGTA AAGTTAGCAT AGATAATTAA TTATTATGA
CACAGTAAC CGAGAATCAT CTGAAAGTGA ATCTTTCATC TGATTGAAAG TATCATGCT GATAACGGAAA
ATCAGAAAA CAACCTTAC TTCGTTAGAAG TAAATT

(SEQ ID NO 117)

Figure 43

TAAGGAAAG GAAACCTGTG AGTTTTCGTT CTTCTCTGTT TGTCAGGTG TTACTTCTCT
GTATGTTGT TCTTGAAAAA CTAGATAAGA AAGTTAGTAA AGTTAGCATA AGTAGTGTAA CTATTATGA
CACAAAGTAAC CGAGAAATCAT CTGAAAGTGA ATCTTTCATC TAATTGACG TATCATGGCT GATACAGACA
ATTAGAAAAA CAACCTTTAC TTCGACCAAG TAAATT

(SEQ ID NO 118)

Figure 44

GGCCTATAGC TCAGCTGGTT AGAGGGCACG CCTGATAAGC GTGAGGTGGA TGGTTCGAGT CCATTAGGC
CACTTTTC TTCTCTGACAG AGAAAACACT GTATAACCTA TTTAAGGGGC CTTAGCTCAG CTGGGAGGC
GCCTGCTTTC CACGCCAGGAG GTCAGGGTT CGATCCCGCT AGGCTCCACC AAAATTGTC TTTGAAACT
AGATAAGAAA GTTAGTAAAG TAGGCATAAA TAGGTAACTA TTATGACAC AAGTAACCGA GAATCATCTG
AAAGTGAATC TTTCATCTGA TTGGAAGTAT CATGGCTGAT ACGAAAAATC AGAAAAACAA CCTTTACTTC
ATCGAAGTAA ATT

(SEQ ID NO 119)

Figure 45

TAAGGAAAG GAAACCTGTG AGTTTCGTT CTTCTTATT TGTCAGTT TGAGAGTTA CTCTCTTTA
TGTCAAGATAA AGTATGCAAG GCACTATGCT TGAAGCATCG CGCCACTACA TTTTGACGG GCCTATAAGCT
CAGCTGGTTA GAGGCCACGC CTGATAAGCG TGAGGTGGAT GGTTCGAGTC CATTAGGCC CACTTTTTCT
TTCTGACATA AGAAATACAA ATAATCATAC CCTTTACGG GGCCTTAGCT CAGCTGGAG AGGGCCCTGCT
TTGCACGCAG GAGGTAGCG GTTCGATCCC GCTAGGCTCC ACCAAATTG TTCTTTGAAA ACTAGATAAG
AAAGTTAGTA AAGTTAGCAT AGATAATTAA TTATTATGAA CACAAGTAAC CGAGAATCAT CTGAAAGTGA
ATCTTTCATC TGATTGAAAG TATCATGGCT GATAAGGAAA ATCAGAAAAA CAACCTTAC TTCGTAGAAG
TAAATT

(SEQ ID NO 120)

Figure 46

TAAGGAAAG GAAACCTGTN AGTTTNCGTN CTTCTCTGTT TGTCAGTTTGTT TCA
TGTCAAGATAA AGTACCGACG GCACGTTGCC TTGGGCAAAG AGCCACTACA TTATTGACGG GCCTATAGCT
CAGCTGTTA GAGGGCACGC CTGATAAGCG TGAGGTCGAT GGTTCGAGTC CATTAGGC CACTTTTCT
TTCTGACAGA AGAAATCATT TGCACATCCT ATTAATAAGG GNCCCTAGCT CAGCTGGAG AGGGCCCTGCT
TTGCACCCAG GAGGTCAAGCG GTTCGATCCC GCTAGGCTCC ACCCAAATT GTTCTTGAA AACTAGATAA
GAAAGTTAGT AAAGTTAGCA TAAGTAGTAT AACTATTAT GACACAAGTA ACCGAGAATC ATCTGAAAGT
GAATCTTCA TCTAATTGCA CGTATCATCG CTGATACAGA CAATTNGAAA AACAAACCTT ACTTCGACGA
AGTAAATT

(SEQ ID NO 121)

Figure 47

TAAGGATAAG GATAACTGTC TAGGACGGT TTGACTAGGT TGGCAAGCG TTTTTTAAT CTTGTATTCT
ATTCCCTTTG CATTGTAAG CGTTGTTCC AAAACATTAA GTTACGATC AAGTATGTTA TGTAATAAT
ATGGTAACAA GTAAATTCAC ATATAATAAT AGACGTTAA GAATATATGT CTTAGGTGA TGTAACTTG
CATGGATCAA TAATTACA

(SEQ ID NO 122)

Figure 48

TAAGGATAAG GAAAGAAGCCT GAGAAGGCTT CTGACTAGGT TGGGCAAGCA TTTATATGTA AGAGCAAGCA
TTCTTATTCA TTTGTGTTGT TAAGAGTACG GTGGTGAGGA CGAGACATAT AGTTTGTGAT CAAGTATGTT
ATTGTAAGA ATAATCATG GTAACAAGTA TATTCACGC ATAATAATAG ACGTTTAAGA GTATTGTCT
TTTAGGTGAA GTGCTTGCAT GGATCTATAG AAATTACA

(SEQ ID NO 123)

Figure 49

CAAATGGAGT TTTTATTCTT TATTATCCTT AACACCCAT TAATTTCCTT GGTGTTAAA CCCAAATCAA
TGTTTGGTCT CACAACTAAC ACATTTGGTC AGTTTGATC CAGTTCTGAA AGAATGTTT TGAACAGTTTC
TTTCAAAACT GAAAACGACA ATCTTTCTAG TTCCCAAAAT AAATAACCAA GGATCAATAC ATAAGTTAC
TAAGGGCTTA TGTT

(SEQ ID NO 124)

Figure 50

CTAATGAAGT TTTTACTTT TTCTTTTCAT CTTTAATAAA GATAAATACT AACAAACA TCAAATCCA
TTTATTATC GGTGGTAAT TAAACCCAAA TCCCTGTTG GTCTCACAAAC TAACATATT GGTCAAGATTG
TATCCAGTTC TGAAGAACCA TTTCCGGTTTC TTTCAAAACT GAAAAGGACA ATCTTTCTAG TTCCAATAAA
ATACCAAAAGG ATCAATACAA TAAGTTACTA AGGGCTATG GT

(SEQ ID NO 125)

Figure 51

AACGAAAGAT TGACCGATTGG TAAGAATCCA CAACAAGTTG TTCTTCATAG ATGTATCTGA GGGTCTGTAG
CTCAGTGGT TAGAGCACAC GCTTGATAAG CGTGGGTCA CAAGTTCAAG TCTTGTCAAG CCCACCATGA
CTTTGACTGG TTGAAGTTAT AGATAAAAGA TACATGATTG ATGATGTAAG CTGGGGACTT AGCTTAGTGTG
GTAGAGGCC TGCTTGGCAC GCAGGGGTC AGGAGTTCGA CTCTCCCTAGT CTCCACCAGA ACTTAAGATA
AGTTCGGATT ACAGAAATTAA GTAAATAAAG ATTGAGATCT TGTTTATTA ACTTCTGTGA TTTCATATTC
ACGGTAATTA GTGTGATCTG ACGAAGACAC ATTAACCTCAT TAACAGATTG GAAAATTGA GTCTGAAATA
AATTGTCAC TCAAGAGTT AGGTTAAGCA ATTAATCTAG ATGAATATGAG AACTAGAAA TAAACTGAAT
CAAGCGTTT GGTATGGAA TTTAGATTGA AGCTGTACAG TGCTTAAGTG CACAGTGCTC TAAACTGAAA
TGGTGAAGTT ACTAACTTGT AGGTAACATC GACTGTTGG GGTGTAT

(SEQ ID NO 126)

Figure 52

AACGAAAGAT TGACGGATGG TAAGAAATCCA CGACAAGGTTG TTCTTCATAG ATGTATCTGA GGGTCTGTAG
CTCAGTTGGT TAGAGCACAC GCTTGATAAG CGTGGGGTCA CAAGTTCAAG TCTTGTCAAG CCCACCATGA
CTTTGACTGG TTGAAGTTAT AGAAAAGAAG ATACATAACT GATGATGTAA GCTGGGGACT TAGCTTAGTT
GGTAGAGCGGC CTGGCTTGGCA CGCAGGAGGT CAGGAGTTCG ACTCTCCTAG TCTCCACCA

(SEQ ID NO 127)

Figure 53

AACGAAAGAT TGATGGCCGG TAAGAATCCA CAAACAAGTTG TTCTTCGAAG ATGTATCTGA GGGTCTGTAG
CTCAGTTGGT TAGAGCACAC GCTTGATAAG CGTGGGGTCA CAAGTTCAAG TCTTGTCAAG CCCACCAAAT
CTGAAAGATA TGTCGTTCAT TATGATTAAA GCTGGGGACT TAGCTTAGTT GGTAGAGGCC CTGCTTTGCA
CGCAGGAGGT CAGGAGTTCG ACTCTCTAG TCTCCACCA

(SEQ ID NO 128)

Figure 54

AACGAAAGAT TGACCGATTGG TAAGAATCCA CAACAAGTTG TTCTTCAATGA CGATGTATCT GAGGGTCTGT
AGCTCAGTTG GTTAGAGGCAC ACGGCTTGATA AGCGTGGGT CACAAGTTCA AGTCTTGTCA GACCCACCAA
ATCTGACTAA CAAGCATTAT TAAATGCTGA ATACAGAAAA ACAGAGACAT TGACTTATG ATAAGCTGGG
GACTTAGCTT AGTTGGTACA GCGGCCTGCTT TGGCACGGAGG AGGTCAAGGAG TTCGACTCTC CTAGTCTCCA
CCA

(SEQ ID NO 129)

Figure 55

AACGAAAGAT TGGTGACCGG TAAGAATCCA CAACAAGTTG TTCTTCGAAG ATGTATCTGA GGGTCTGTAG
CTCAGTTGGT TAGAGCCAC GCTTGATAAG CGTGGGTCA CAAGTTCAAG TCTTGTCAAG CCCACCACTA
CTGACCGAAGT GATGAATAAT CACAAGCTGC TAGATGAAA GATATGTCGT TCATTATGAT TAAAGCTGGG
GACTTAGCTT AGTTGGTAGA GGCCTGCTT TGGCACGGAG AGGTCACTC CTCAGTCTTC CTA
CCA

(SEQ ID NO 130)

Figure 56

TAAGGAAAGAT CGAGAATTGG AAAGAGGGTCG GATTATCCG GATGATCCTT CTCCCATCCTTA TTAGAACATA
 GATCGCAGGC CAGTCAGCCT GACGATCGCT TGCAGGGCGTG CCGCCTTCGT TTCTCTTTCT TCATTTGTTGA
 TTGCTCACGG GCCGTAACCGC AGCTGACCGCT GCTGGCCCTG CGCAGGGCGG GCCCATCAGG GCCGACGGCC
 GGTGGGCCCTT GCNAAGCTTC GCTTCCGGGT GGATCTGTGG ATCGGGTAGT AGCGTTGTGCG TCGGTATCTG
 GGCTTGTAGC TCAGTTGGTT AGAGCCACACG CTTGATAAGC GTGGGGTCTGG AGGTTCAGT CCTCCCCAGGC
 CCACCAAGTT ACTTGATGAG GGGCCGTAGC TCAGGTGGGA GAGGCACCTGC TTGCAAGCA GGGGGTGTGTC
 GGTTCGATCC CGTCCGGCTC CACCATCATG TTGGTGTGTA GACGGATATT GGCAATCAAC AAAAGAAAGA
 AACAAAGTTG CGGACTNTTA CGAAAGTCTG CCTGGTCTGT ATGAAATCGT GAAGAGAAAGA TGTAATCGGA
 TCAACTGAAG AGTTGATGTC GCAAGAAAGCT TGCTCAAGCC TTGCATAATG ATTGATGTGT TAAACCGCCA
 TCACCCGATTG TATCTCGAGA AGCTGGTCTT TCTGCTGATA CTGTTGAAAC GAGGATTTGC AGTCGAATGG
 CAACATTGG CGTCGCATAA TGGGGCTTAA AGAGCTGAGT TTGATGGAT ATTGGCAATG AGAGTGTATCA
 AGTGTCTTAA GGGCATTGGT GGATGCCCTTG GCATGGCAC

(SEQ ID NO 131)

Figure 57

TAAGGGAGGAT CGAGAATTGG AAAGAGGGCG GATTATCCG GATGATCCTT CTCCATCTTA TTAGAACATA
GATCGCAGNC CAGTCAGCCT GACCGATCGCT TGCAGGGGTG CGCCTCTCG TTCTCTTCT TCATTGTTGA
TTGCTCACGG GCCGTACCGC AGCTGACGCT GCTGGCCCTG GCTGGGGCTG GNCCATCAGG GCCGACGGCC
GGTCGGCCTT GCGAAGCTTC GCTTCGGGGT GGATCTGTGG ATCGCGTAGT AGCGTTGTGG TCGGTATCTG
GGCTTGTAGC TCAGTTGGTT AGAGCCACACG CTTGATAAGC GTGGGGTGG AGGTTCAAAT CCTCCAGGC
CCACCAAGTT ACTTGATGAG GGGCCGTAGC TCAGCTGGGA GAGGCCACTGC TTTGCAAGCA GGGGGTGTGTC
GGTTCGATCC CGTCCGGCTC CACCATCATG TTGGTGTGA GACGGATT GGCAATCAAC AAAAGAAAGA
AACAAAGTTG CGGACTNTTA CGAAAGTCTG CCTGTTCTGT ATGAAATCGT GAAGAGAAGA TGTAAATGGGA
TCAACTGAAG AGTTGATGTC GCAAGGAAGCT TGCTCAAGCC TTGCTAAATG ATTGATGTGT TTAACGGCCA
TCACCGATTG TATCTCGAGA AGCTGGTCTT TCTGCTGATA CTGTTGAAAC GAGGCATTGC AGTCGAATGG
CAACATTGG CGTCGCATAA TGCGGCTTA AGAGCTGAGT TTTGATGGAT ATTGGCAATG AGAGTGTATCA
AGTGTCTTAA GGGCATTGGT GGATGCCCTTG GCATGGCAC

(SEQ ID NO 132)

Figure 58

CCTTAAGAA CTGTTCTTTCAGTGCTCAC ACAGATTTGTC TGATGAAAAG TAAATAGCAA GGGGTCTTGC
GAAGCAGACT GATAACCTCCC CTTCGTCTAG AGGCCAGGA CACCGCCCTT TCACGGGGT AACAGGGGT
CGAATCCCCT AGGGGAGGCC ACTTGGCCGG TAATGTGTGA AAGCGTTGCC ATCAGTATCT CAAAACGTGAC
TTACGAGTCA CGTTTGAGAT ATTTGCTCTT TAAAATCTG GATCAAGCTG AAAATTGAAA CACAGAACAA
CGAAAGTTGT TCGTGAGTCT CTCAAATTTCGCAACACGA TGATGAATCG TAAGAACAT CTTGGGTTG
TGA

(SEQ ID NO 133)

Figure 59

CCTTAAGAA CTGTTCTTTCAGTGCTCAC ACAGATTGTC TGATGAAAAA CGAGCAGTAA AACCTCTACA
GGCTTGTAGC TCAGGGGGTT AGAGGGCACC CCTGATAAGG GTGAGGGTCGG TGTTCAAGT CCACTCAGGC
CTACCAAATT TTCCCTGAAT ACTGCCGTGT GAAATAACTC ACATAACTGAT GTATGCCCG TTATTCCACG
CCTTGTCTCA GGAAAATTAA TCGGTAAGA GTTCTGTACT ACACGATGGG GCTATAGCTC AGCTGGAGA
GGGCCCTGCTT TGCACGGCAGG AGGTCTGGG TTTCGATCCCG CATAAGCTCCA CCATATCGTG AGTGTGTTACG
AAAAAATACT TCAGAGTGTAA CCTGAAAGGG TTCACTGGCA AGTTTGGCTC TTTAAAATC TGGATCAAGC
TGAAAATGAA AACACAGAAC AACGAAAGTT GTTCGTGAGT CTCCTCAAATT TTCGCAACAC GATGATGAAT
CGTAAGAAC ATCTTCGGGT TGTGA

(SEQ ID NO 134)

Figure 60

CCTTAAAGAA GCGTACTTTG CAGTGCTCAC ACAGATTGTC TGATGAAAAG TAAATAGCAA GGGGTCTTGC
GAAGGAGACT GATAAGTCCC CTTCGTCTAG AGGCCAGGA CACCGCCCTT TCACGGGGGT AACAGGGGTT
CGAATCCCCT AGGGGAGGCC ACTTGGGGGG TAATGTGTGA AAGCGTTGCC ATCAGTATCT CAAAACGTGAC
TTACGAGTCA CGTTTGAGAT ATTTGCTCTT TAAAATCTG GATCAAGCTG AAAATTGAAA CACAGAACAA
CGAAAGTTGT TCGTGAGTCT CTCAAATTTC CGCAACACGA TGATGAATCG TAAGAAACAT CTTCGGGTTG
TGA

(SEQ ID NO 135)

Figure 61

CCTTAAAGAA CTGTTCTTTC AAGTGCTCAC ACAGATTGTC TGATGAAAAA CGAGCAGTAA AACCTCTACA
GGCTTGTAGC TCAGGGTGGTT AGAGGCCACC CCTGATAAGG GTGAGGGCGG TGGTTCAAGT CCACTCAGGC
CTACCAAATT TTCCCTGAAT ACTGGCTTGT GAAATAACTC ACATACTGAT GTATGCTTCG TTATTCACCG
CCTTGTCTCA GGAAAATTAA TCGGTAAAGA GGTTCTGACT ACACGATGGG GCTATAGCTC AGCTGGAGA
GCGCCCTGCTT TGCACGGCAGG AGGTCTGGGG TTTCGATCCCG CATAAGCTCCA CCATCTCGTG AGTGTGTTACG
AAAAAATACT TCAGAGTGTAA CCTGAAAGGG TTCAC TGCGA AGTTTTGCTC TTAAAAAATC TGGATCAAGC
TGAAAATTGA AACACAGAAC AACGAAACTT GTTCGTGAAGT CTCTCAAATT TTTCGCAACAC G

(SEQ ID NO 136)

Figure 62

CCTTAAGAA GCGTACTTTG AAGTGTCTCAC ACAGATTGTC TGATGAAAAG TGAATAGCAA GGGGTCTTGC
GATTGAGACT TCAGTGTCCC CTTCGTCTAG AGGCCAGGA CACCGCCCTT TCACGGGGGT AACAGGGGT
CGAATCCCCT AGGGGACGCC AGCGTCAAA CTGATGAGGT CAAACCTCCA GGGACCCAC TTGCTGGTT
GTGAGTGAAA GTCACCTGCC TTAATATCTC AAAACTGACT TAGGAGTCAC GTTTGAGATA TTTGCTCTTT
AAAAATCTGG ATCAAGGCTGA AAATTGAAAC ACAGAACAAAC GAAAGTTGTT CGTGAGTCTC TCAAATTTTC
GCAAACACGAT GATGAATCGT AAGAAAACATC TTCGGGGTTGT GA

(SEQ ID NO 137)

Figure 63

CCTTAAGAA ACGGTCTTTC AAGTGCTCAC ACAGATTGTC TGATGAAAAA CGAGCAGTAA AACCTCTACA
GGCTTGTAGC TCAGGGGGTT AGAGGGCACC CCTGATAAGG GTGAGGGTGG TGTTTCAAGT CCACTCAGGC
CTACCAAATT TTCCCTGAAT ACTGGCTTGT GAAATAACTC ACATACTGAT GTATGCTTCG TTATTCCACG
CCTTGTCTCA GGAAAATTAA TCGGTAAGA GGTTCGTGACT ACACGATGGG GCTATAGCTC AGCTGGGAGA
GCGCCCTGCTT TGCACGCCAGG AGGTCTGGG TTCGATCCCG CATAAGCTCCA CCATCTCGTG AGTGTGTTACG
AAAAAATACT TCAGACTGTA CCTGAAAGGG TTCACTGGGA AGTTTTGCTC TTAAAAAATC TGGATCAAGC
TGAAATTGA AACACAGAAC AACGAAAGTT GTTCGTGAGT CTCTCAATT TTGGCAACAC GATGATGAAT
CGTAAGAAC ATCTTGGGT TGTGA

(SEQ ID NO 138)

Figure 64

CTAAGGGATAT ATTGGAAACA TCTTCCTTCGG AAGATGGGA ATAACGTGAC ATATTGTATT CAGTTTGAA
TGTTTATTAA ACATTCAAAT ATTTTTGGT TAAAGTGATA TTGCTTTGAA AAATAAAGCA GTATGGGAGC
GCTTGACTAA AAAAATTGT ACATTGAAAA CTAGATAAGT AAGTAAATA TAGATTTAC CAAGCAAAAC
CGAGTGAATA AAGAGTTTA AATAAGCTTG AATTCTAAAG AAATAATCGC TAGTGGTCGA AGAACACACTC
ACAAGATTAA TAACGGGTTT AAATCTTT ATAACAGAAC GTAACCTTCAT GTTAACGTTT GACTTATAAA
AATGGGGAA ACATA

(SEQ ID NO 139)

Figure 65

CTAAGGATAT ATTCGGAACA TCTTCTTCAG AAGATGGGA ATAACGTGAC ATATTGTATT CAGTTTGAA
TGTTTATTAA ACATCCAAT ATTTTTGGT TAAAGTGATA TTGCTTATGC GAGCNCCTGA CAATCTATT
TTTTTAAGA AAGGGTTGT CAGACAATGC ATTAAGAAAA ATTAAGGGG AGTTTACTTT TGTAATGAG
CATTTGATT TTGAAAATA AAGCACTATG CGAGCGCTTG ACTAAAAAGA AATTGACAT TGAAAACTAG
ATAAGTAAGT AAAATATAGA TTTTACCAAG CAAACCGAG TGAATAAGA GTTTAAATA AGCTTGAAATT
CATAAGAAAT ATCGCTAGT GTTCGAAAGA ACACTCACAA GATTAATAAC GGGTTAAAT CTTTTATAA
AGAAAACGT TTAGCAGACA ATGAGTAA TTATTTAA GCAGAGTTA CTTATGTAAA TGAGCATTAA
AAATAATGAA AACGAAGCCG TATGTGAGCA TTTGACTTAT AAAATGGTG GAAACATA

(SEQ ID NO 140)

Figure 66

CTAAGGATAT ATTCGGAACA TCTTCTTCAG AAGATGCCGA ATAACGTGAC ATATTGATT CAGTTTGAA
TGGTTATTA ACATTCAAAT ATTTCCTGGT TAAAGTGTATA TTGCTTATGC GAGCGCTTGA CAATCTATT
TTTTAAAGA AAGGGTTGT CAGACAATGC ATTAAGAAAA ATAAAGCGG AGTTTACTTT TGAAATGAG
CATTGATT TTGAAAATA AAGCACTATG CGAGCGCTTG ACTAAANGA AATTGTACAT TGAAAACTAG
ATAAGTAAGT AAAATATAGA TTTTACCAAG CAAACCGAG TGAATAAGA GTTTGAAATA AGCTTGAAATT
CATAAAGAAAT ATCGCTAGT GTTCGAAAGA ACACTCACAA GATTAATAAC GCGTTTAAAT CTTTTTATAAA
AGAACCGTAA CTTCATGTT ACGTTTGACT TATAAAATG GTGGAAACAT A

(SEQ ID NO 141)

Figure 67

CTAAGGATAT ATTGGAAACA TCTTCTTCAG AAGATGGGA ATAACGTGAC ATATTGATT CAGNTTGAA
TGTTTATTTA ACATTCAAAA AATGGGCCTA TAGCTCAGCT GGTTAGAGCG CACGCCCTGAT AAGCGTGAGG
TCGGTGGTTC GAGTCCACTT AGGCCACCA TTATTGTAC ATTGAAACT AGATAAGTAA GTAAAATATA
GATTTACCA AGCAAAACCG AGTGAATAAA GAGTTTAAA TAAGCTTGAA TTCATAAGAA ATAATCGCTA
GTGTTCGAAA GAACACTCAC AAGATTAATA ACGCGTTAA ATCTTTTAT AAAAGAACGT AACTTCATGT
TAACGTTTGA CTTATAAAA TGGTGGAAC ATA

(SEQ ID NO 142)

Figure 68

CTAAGGATAT ATTCGGAACA TCTTCYTCAG AAGATCGGAA ATAATGTGAC ATATTGTATT CAGTTTGAA
TGTTTATTAA ACATTCAAAT ATTTTTGGT TAAAGTGATA TTGCTTATGC GAGGGCTTGA CTAAAAAGAA
ATTGTCACATT GAAAACCTAGA TAAGTAAGTA AAANTATAGA TTTTACCAAG CAAAACCGAG TGAATAAAGA
GTTTTAAATA AGCTTGAATT CATAAGGAAT AATCGCTAGT GTTCGAAAGA ACACTCACAA GATTAAAAC
CGGTTAAAT CTTTATAA AAGAACGTA CTTCATGTT ACGTTTGAAT TATAAAATG GTGGAAACAT
A

(SEQ ID NO 143)

Figure 69

CTAAGGATAT ATTGGAAACA TCTTCTACGA AGATGAGGGA ATAACGTCGAC ATATTGGTATT CAGTTTGAA
TGTTTATTAA CATTCAATTG TACATTGAAA ACTAGATAAG TAAGTAAGAT TTTACCAAGC AAAACCCAGT
GAATAGAGTT TAAATAAGC TTGAATTCAAT AAATAATCGC TAGTGTTCGA AAGACNTCCA CAAGATTAAAT
AACTAGTTT AGCTATTAT TTTGAATAAC AATTCAAAT ATGGTGGGAC ATA

(SEQ ID NO 144)

Figure 70

AAGGATAAGG AACTGCACAT TGGTCTGTGTT TAGTCTGTGAG AGGTCTGTG GGGCCTTAGC TCAGGCTGGGA
GAGCCCTGCC TTGGCACGCA GGAGGTCAAGC GGTTCGATCC CGCTAGGCTC CATTGGTGAG AGATCACCAA
GTAATGCACA TTGAAATTG AATATCTATA TCAAATAGTA ACAAGAAAAT AAACCGAAAA CGCTGTAGTA
TTAATAAAGA GTTTATGACT GAAAGGTCAA AAAATAA

(SEQ ID NO 145)

Figure 71

AAGGAAATGG AACACGTTA TCGTCTTATT TAGTTTGAG AGGTCTTGTG GGGCCTTAGC TCAGCTGGGA
GAGGCCCTGC TTNGCACGCA GGAGGTCAGC GGTTCGATCC CGCTAGGCTC CATCAGGATA CANTCCTACT
AAACTAATA CAAGTGAAGT TGAACACGGCA ACTCACTTCC TAGGAAAATA GACAATCTTC GCTTGTGTC
AAGGCACACA TGGTCAGATT CCTAATTTC TACAGAAGTT TCGCTAAAGC GAGCGTTGCT TAGTATCCTA
TATAATAGTC CATNGAAAT TGAATATCTA TATCAAATT CACGATCTAG AAATAGATTG TGCAAAACGTA
ACAAGAAATT AACCCGNAAA CGCTG

(SEQ ID NO 146)

Figure 72

AAGGATAAGG AACTGCACAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGAG GGGCCTTAGC TCAGCTGGGA
GAGCGCCCTGCG TTTGCCACGCA GGAGGTCAGC GTTTCGATCC CGCTAGGCTC CATTGGTGAG AGATCACCAA
GTAATGCACA TTGAAATAATG AATATCTATA TCAAATAGTA ACAAGAAAAT AAACCGAAC GCTGTAGTAT
TAAAAGAGTT TATGACTGAA AGGTCAAGAAA ATAA

(SEQ ID NO 147)

Figure 73

CTAAGGATAT ATTGGAAACA TCTTCTTACG AAGATGCAGG AATAACATTG ACATATTGTA TTCAGGNTGTG
AATGGCTCATT GGAGNATTCA TNGCATNATT TGGTNCATTG ACANCTAGAT AAGNAAGTAA AATTATGAT
TTTACCAAGC AAAACCGAGT GAATTAGAGT TNTNNAACAA GCTTTGATT CAAAAGAAA TAATCGCTAG
TGTTCGAAAG AACACTCAC A GATTANTAAC ATCTTGGTT TTCAACCCGAC TTGTTCGTNT CGAAAGTCAA
AAAA

(SEQ ID NO 148)

Figure 74

AAGGATAAGG AACTGGCAT TGGTCTTGTG TAGTCTTGAG AGGTCTTGTG GGGCCTTAGC TCAGCTGGGA
GAGGCCCTGC TTGGCACCGCA GGAGGTCAAGC GGTTGGATCC CGCTAGGCTC CATTGGTGAG AGATCACCAA
GTAATGCCACA TTGAATAATTG AATATCTATA TCAAATAGTA ACAAGAAAT AAACCGAAAA CGCTGTAGTA
TTAATAAGAG TTATGACTG AAAGGTCAAA AAATAA

(SEQ ID NO 149)

Figure 75

AAGGATAAGG AACTGGCAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGTG GGGCCTTAGC TCAGCTGGGA
GAGGCCCTGC TTTGCACGCA GGAGGTCA GC GGTTCGATCC CGCTAGGCTC CATTGGTGA AGATCACAA
GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA ACAAGAAAAT AACACGAAAA CGCTGTAGTA
TTAATAAGAG TTTATGACTG AAAGGTCA GA AAAATAA

(SEQ ID NO 150)

Figure 76

AAGGAAAGG AACTGGCAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGAG GGGCCTTAGC TCAGCTGGGA
GAGCGCCTGC TTTGCACGCA GGAGGTCAAGC GTTTCGATCC CGCTAGGCTC CATTGGTGAG AGATCACCAA
GTAATGCCACA TTGAAAAATTG AATATCTATA TCAAATAGTA ACAAGAAAAT AAACCGAAAAA CGCTGTAGTA
TTAATAAGAG TTTATGACTG AAAGGTCAAG AAAATAA

(SEQ ID NO 151)

Figure 77

AAGGATAAGG AACTGGCAT TGGTCTTGTT TAGTCTTGAG AGGTCTTAGC GGGCCTAGC TCAGCTGGGA
GAGGCCCTGC TTTGCACGCA GGAGGTCAAGC GGTTCGATCC CGCTAGGCTC CATTGGTGAAG AGATCACCAA
GTAATGCACA TTGAAAATTG AATATCTATA TCAAATAGTA ACAAGAAAT AAACCGAAC GCTGTAGTAT
TAAAAGAGTT TATGACTGAA AGGTCAAGAAA ATAA

(SEQ ID NO 152)

Figure 78

AAGGATAAGG AACTGGCAT TGGTCTTGTT TAGTCTTGAG AGGTCTTGAG GGGCCTTAGC TCAGCTGGGA
GAGGCCCTGC TTGCAACGCA GGAGGTCAAGC GGTTCGATCC CGCTAGGCTC CATTGGTGAAG AGATCACCAA
GTAATGCACA TTGAAAAATTG AATATCTATA TCAAATAGTA ACAAGAAAAT AACCGAAC GCTGTAGTAT
TAAAAGAGTT TATGACTGAA AGGTCAAAAA TAA

(SEQ ID NO 153)

Figure 79

TAAGGAAGAT CGAGAATTGG AAAGAGGTG GATTATCCG GATGATCCTT CTCCATCTTA TTAGAACATA
 GATCGCAGGC CAGTCAGCCT GACGGATCGCT TGCAGGGCTG CCGCCTTCGT TTCTCTTCT TCATTGTTGA
 TTGCTCACGG GCCGTACCGC AGCTGACGCT GCTGGCCCTG CGCAGGGCG GCCCACAGG GCCGAACGGC
 CGGTGCGCCT TGCNAAGCTT CGCTTCGGGG TGGATCTGTG GATCGCGTAG TAGCGTTGCT GTCGGGTATCT
 GGGCTGTAG CTCAGTTGGT GCTTGATAAG CGTGGGTCTG GAGGTTCAAG TCCTCCAGG
 CCCACCAAGT TACTTGATGA GGCGCCGTAG CTCAGCTGGG AGAGCACCTG CTTTGCAGG AGGGGGTCTG
 CGGTTCGATC CCGTCGGCT CCACCATCAT GTTGGTGTG AGACGGATAT TGCCAATCAA CAAAGAAAG
 AAACAAGTT GCGGACTNTT ACGAAAGTCT GCCTGTTCTG TATGAAATCG TGAAGAGAAG ATGTAATCGG
 ATCAACTGAA GAGTTGATGT CGCAAGAAC TTGCTCAAGC CTTGCATAAT GATTGATGTG TTTAACGCC
 ATCACCGATT GTATCTCGAG AAGCTGGTCT TTCTGCTGAT ACTGTTGAA CGAGCATTG CAGTCGAATG
 GCAAACATTG GCGTGCATA ATGCGGCTT AAGAGCTGAG TTTTGATGGA TATTGGCAAT GAGAGTGATC
 AAGTGTCTTA AGGGCATTGG TGGATGCCCTT GGCAATGCAC

(SEQ ID NO 154)

Figure 80

AAGGAGCACC ACCGAAACA CTCCAATTGG TGGGGTGTAA GCCGTCAGGG GTTCTCGTCT GTAGTGGACG
GAAGCCGGGT GCACAAAC AAGCAAGCCA GACACACTAT TGGGTCTGA GGCAACATCT CTGTTGGTT
CGGGATGTTG TCCCACCATC TTGGTGGTGG GGTGTGGTGT TTGAGAATTG GATACTGGTT GCGAGCATCA
ATTGGATGCG CTGCCCTTTG GTGGCCGTGTT CTGTTGTGCA ATTATTATTCT TTGGTTTTTG TGTTAT

(SEQ ID NO 157)

Figure 81

AAGGAGGCC ACCGAGAAACA CCCCAATTGG TGGGGTGTGA GCGGTGAGG GTTCTCGTCT GTAGTGGACG
AGGGCCGGGT GCACAAACAAC AGGCAATCGC CGAACACACT ATTGGCCCT GAGACAACAC TCGGCCGACT
GAGGTGGACG TGGTGTCCCT CCATCTTGTT GGTGGGTGT GGTGTTGAG CATTGAATAG TGGGTTGCGAG
CATCTAGCCG GATGCGTTCC CCAGTGGTGC GCGTTCGTCA AAAATGTGTA ATTTTCTTT TGGTTTTGT
GTTCGT

(SEQ ID NO 158)

Figure 82

AAGGAGGCC ACCGAGAAACA CCCCAAATTGG TGGGGTGTGA GCGGTGAGGG GTTCTCGTCT GTAGTCGACG
AGGGCCGGGT GCACAAACAAC AGGCAATCGC CGGACACACT ATTGGGCCCT GAGACAAACAC TCGGCCGACT
GAGGTGAGCG TGCTGTCCCT CCATCTTGGT GGTGGGGTGT GGTGTTGAG CATTGAATAG TGTTTGCGAG
CATCTAGACG GATGGCTTCC CCAGTGGTGC GCGTTCGTCA AAAATGTGTA ATTTTCTTT TGTTTTGT
GTTCGT

(SEQ ID NO 159)

Figure 83

AAGGAGGCACC ACGAGAAACA CCCCAATTGG TGGGGTGTGA GCCGGTGGGG GTTCTCGTCT GTAGTGGACG
AGGNNDNGGGT NNACAAACAC NGCCAATCGC CGGACACACT ATTGGCNCT GAGACAAACAC TCGGCCGACT
GAGGTGACG TGGTGTCCCT CCATCTTGCT GGTGGGGTGT GGTGGTTGAG CATTGAATAG TGTTTGGAG
CATCTAGCCG GATGGTTCC CCAGTGGTGC CGCTTCGTCA AAAATGTGTA ATTTTCTNT TGGTTTTGT
GTTCGT

(SEQ ID NO 160)

Figure 84

AAGGAGGCC ACCGAGAAACA CTCCAATTGG TGGAGGTGTGA GCCGGTGGAGG GTAGCTCGTCT GTAGTGGACG
AGGGCCGGGT GCACAAACAGC AGACAATCGC CAGACACACT ATTGGGCCCT GAGACAAACAC TCGGCCGACT
TTGGTGTGAGC TGGTGTCCCT CCATCTTGCT GGTGGGGTGT GGTGTGTGAG CATTGAATAG TGTTTGCAG
CATCTAGACG GATGGGTGTC CCTCGGGCCG CGTGTCGTC AAAAATGTGT AATTTTTCT TTGGTTTT
GTGTTGCT

(SEQ ID NO 161)

Figure 85

AAGGAGGCACC ACGAGAAACA CTCCAATTGG TGGAGTGTGA GCCGTGAGGG GTTCTCGTCT GTAGTGGACG
GGAGCCGGGT GCACAAACAA AGGCAATCGC CAGACACACT ATTGGCCCT GAGACAAAC AC TCGGCCGCT
TTGAGTCGAA TGCGTGTCCC TCCATCTTGG TGCGTGGGTG TGGTGTGTGA GCATTGAATA GTGGTTGCGA
GCATCTAGAC GGATGGCTTG CCTTCGGGCC GCGTGTTCGT CAAAATGTG TAATTTCGTTT TTTTGGTTT
TGTGTTCGT

(SEQ ID NO 162)

Figure 86

AGGGAGGCC AGAACCGCAT CCCGGCTGGG GTGTGGTTTC GGGGTGGTGT CGAGGGTGGNC
GGCAGCAGGC AGTAACCNCC GGAACACTGT TGGGTGGTGA GNNAACACCC GTGGTGGTGT TGTGCTCCCC
GTGGTGNCGG GGTGGTGGTGT TTGAGTGGTT GATAGTGGTT GCGAGCATCT GGCAAAAGACT GTGGTAAAGCG
GTTTTTGTGG ANTGTGGTCT GGTGGTGTGT

(SEQ ID NO 163)

Figure 87

AAGGAGCACC ACGAGAAACA CTCCAATTGG TGGGGTGTGA GCCGTGAGGG GTTCTCGTCT GTAGTGGACG
AGGGNCGGGT GCACAAACAA AGNCAATCGC CAGACACACT ATTGGNCCCT GAGACAAACAC TCGGCCGACT
TNGGTTGAAG TGGTGRCCCT CCATCTTGGT GGTGGGTGT GGTGTTGAG TATTGGATAG TGTTGGAG
CATCTAANTG AACGGGTGCGC CGNCAACGGT TACGTGTTCG TTGTTGTAA TTNTTTCTAT TGTTTTGT
GTTCGT

(SEQ ID NO 164)

Figure 88

AAGGAGCACC ACGAGAAACA CTCCAATTGG TGGGGTGTGA GCCGGTGAGG GTAGTGGACG
AGGGCCGGGT GCACAAACAAAC AGGCAATCGC CAGACACACT ATTGGNCCCT GAGACAAACAC TCGGCCGACT
TTGGTCAAAG TGGTGTCCCC CCATCTTGGT GGTGGGGTGT TATTGGATAG TGGTTGCAGAA
CATCTAAATG AACGGGTTCG CGGCAACGGT TACGTGTTCG TTTTAGTGTAA ATGNTTTCTA ATGGTTTTG
TGTTCGT

(SEQ ID NO 165)

Figure 89

AAGGAGGCC ACCAGAACCTG GGCCGGCCCC GCAGATCGGG GGATCAGCTG AGCTTTCAGG CGATTCTGTTG
GATGGCCTCG CACCTGTAGT GGGTGGGGT CTGGTGCAC T CAACAAACTT GGGGTGGGAT GCGGGAAAGC
ATCTGGGAA AATCATCAGA CACACTATTG GGCTTIGAGA CAACAGCCCC GCAGNCCTGN CCCGTTGGGG
GCAGNGGGTG TGTTGTGCC TCACTTGGT GGTGGGGTG GTGTTTGATT TGTGGATACT GTTGGCGAGC
ATCTAGGGCG CAGAATGTGT GGTCTCACTC CTTGTGGTG GGGCCTGGTT TTGTGTGCGA TTGATGTGCA
ATTTCCTTTG AAACTCAATT TTGGTTTT GTGTTGT

(SEQ ID NO 166)

Figure 90

AAGGAGGCC ACCAAAAACT CCCCAATTGG TGGGGTGTAA GCCGGTGGGG GTTCCCCGTCT GTAGTGGACG
GGGGCCGGGT GCGCAACAGC AAGCGAAACG CCGGACACAC TATGGGTCC TGAGGCAACA CTCGGGTTTG
TCCCCCTCAG GGATTTCTG GGTGTTGTCC CACCATCTG GTGGTGGGGT GTGGTGTG AGAATTGGAT
AGTGGTGGC AGCATCAAAT GGATGCGTTG CCCCTACGGG TAGCGTGTTC TTTGTGCAA TTTTATTCTNT
TGGTTTTTGT GTTTGT

(SEQ ID NO 167)

Figure 91

AAGGAGGCACC ACGAGAAGCA CTCCAACCTGG TGGGGTGCAA GCCGGTGAGGG GTTCTCGTCT GTAGTGGACG
AGAGGCCGGGT GCGCGAAC GAACGAGCCA GACACACTAT TGGGTCTGA GGCAACACTC GGGCTTGCC
AGAGCTGRTG TCCCACCATC TTGGTGGTGT TTGAGAATTG GATACTGGTT GCGAGCATCA
AATGGATGCG TTGCCCCCTAC GGGTGGCGTG TTCTTTGTG CAATTATT CTTTGGTTT TGTGTTGT
(SEQ ID NO 168)

Figure 92

AAGGAGCACC ACGAAAAACA CCCCAACTGG TGGGGTGTAA GCCGTGAGGG GCTCCCGTCT GTAGTAGACG
GGCGCCGGGT GCGCAACAGC AAGCGAACCCA GACACACTAT TGGGTCTGA GGCACAACTC GGGCTGTCT
TGGACTCGTC CAAGAGTGT GTCCCACCAT CTTGGTGGTG GGGTGTGGT TTTGAGAATT GGATAAGTGGT
TGCGAGGCATC ANCTGGATGC GTTGCCCCCA GGGTAGGCGT GTTCTTTGT GCAATTNTAT TCNNTGGTT
TTGTGTTAGT

(SEQ ID NO 169)

Figure 93

AAGGAGCACC ACGAAAAACA CTCCGGATCC GGTGGGGTGT GAGGCCGTGAG GGAGGCCGTG CCTGTAGTGG
GTGTGGTTG GGTGGCGAC AACAAATGGG AAAAATCGCT GGGCACACTA TGGGGCTTTG AGGCAACACC
TGGTTTGTTT TGGGTGGTGT CGCTCCATCT TGCTGGTGGG GTGTGGTGT TGAGTGTGG ATAGTGGTTG
CGAGGCATCTA AGCAAAGCT GTTGTGAC GTTGTGTGTC GGTTTTGTG CAGTGTGTG TGTGT

(SEQ ID NO 170)

Figure 94

AAGGAGGCC ACC AGAAAAACA CTCCAATTGG TGGGGTGTAA GCCGTGAGGG GTAGCTCATCT
AGAGCCCCGGT GCACAAACAGC AAATGAATCG CCAGACAC AC TGTTGGGTCC TGAGGCAACA CTCAGGGCTTG
TCCCATGTG GGCTGTGATCG GGTGCTGTCC CCCCATCTTG GTGGTGGGGT GTGGTGTG AGTATGGAT
AGTGTTGGC AGCATCTAAA TGGATAACGTT GCCAGTAATG GTGGCGTATT CATTGAAAAT GTGTAATT
CTTCTTTGGT TTGTGTGT

(SEQ ID NO 171)

Figure 95

AAGGAGGCACC ACGAAAACA CTCCAATTGG TGGGGTGTAA GCGGTGAGGG GTTCTCATCT GTAGTGGACG
AGAGGCCGGGT GCACAAACAGC AAATGAATCG CCAGAACAC TGTTGGTCC TGAGGAACA CTCAGGCTTG
TCCCATGTTG GGCTTGATCG GGTGCTGTCC CCCCATCTTG GTGGTGGGT GTGGTGTGG AGTATTGGAT
AGTGGGTGCG AGCATCTAAA TGGATACTGTT GCCAGTAATG GTGGCGTGT CATTGAAAAT GTGTAATT
CTTCTTTGGT TTGTTGTT

(SEQ ID NO 172)

Figure 96

AAGGAGCACC ACGAAAAACA CTCCAATTGG TGGGTGTAA GCCGTGAGGG GTTCTCATCT GTAGTGGACG
AGAGCCGGGT GCACAAACAGC AAATGAATCG CCAGACACAC TGTTGGTCC TGAGGCAACA CTCAGGCTTG
TCCCATGTC GGCTTGATCG GGTGCTGTCC CCCCATCTTG GTGGTGGGGT GTGGTGTG AGTATGGAT
AGTGGTGG AGCATCTAAA TGGANACGTT GCCAGTAATG GTGGCGTGT CATGAAAT GTGTAATT
CTTCTTGGT TTGTGTGT

(SEQ ID NO 173)

Figure 97

AAGGAGCACC ATTCTCTCAGT CGAATGAACT GAGAACATAA AGCGAGTATC TGTAGTGGAT ACATGCTTGG
TGAATATGTT TTATAATCC TGTCCACCCC GTGGATAGGT AGTCGGAAA ACGTCGGACT GTCATAAGAA
TTGAAACGGCT GGCACACTGT TGGGT CCTGA GGCAACACAT TGTGTTGTCA CCCTGCTTGG TGTTGGGTG
TGGTCCCTTGA CTTATGGATA GTGGTTGCGA GCATCTAAC ATAGCCTCGC TCGTTTCGA GTGAGGCTGG
TTTTTGCAAT TTTATTAGCT

(SEQ ID NO 174)

Figure 98

CCTAATGATA TGTGATTGCGG TGAAGTGGTC ACACAGATTG TCTGATGAAA AAGTAACGAG CAGAAATAACC
TTTATAGGCT TGTAGCTCAG GTGGTTAGAG CGCACCCCTG ATAAGGGTGA GGTCGGTGGT TCAAGTCCAC
TCAGGGCCCTAC CACTTCCTCGA AGTGGAAAAG GTACTGCACG TGACTGTATG GGGCTATAGC TCAGCTGGGA
GAGGGCCCTGC CTTGCCACCA GGAGGTCAAGC GGTTCGATCC CGCTTAGCTC CACCATATAG TCCTGTATTI
CAATACTTCA GAGTGTACTG GCAACAGTAT GCTGCCAAGT ATTTTGCTCT TTAACAATCT GGAACAAAGCT
GAAAATTGAA ACATGACAGC TGAAACTTAT CCCTCCCGTAG AAGTATTGGG GTAAGGATA ACCTGTCTA
GAGTCTCTCA AATGTAGCAG CACGAAAGTG GAAACACCTT CGGGTTGTGA

(SEQ ID NO 195)

Figure 99

CCTAAATGATA TTGATTCGCG TGAAGTGCTC ACACAGATTG TTTGATAGAA ACGTAATGAG CAAAAGGGCT
ACCTGTGAT GTAATGAGTC ACTGACTCAT GCTGATACGA ACCGATTAAG ACAGTCAGTT TAATCGGATT
TTCGTGCCC CATCGTCTAG AGGCCTAGGA CACTGCCCTT TCACGGCTGT AACAGGGTT CGAATCCCT
TGGGGACGCC ATTCCGATAAT GAGTGAAGA CATTATCACC GGTTCTTGGA ACCGAAAACA TCTTAAGAT
GACTCTGCG AGTCGTGTT AAGATATTGC TCTTTAACAA TCTGAAACAA GCTGAAAATT GAAACATGAC
AGCTGAAACT TATCCCTCCG TAGAAGTATT GGGTAAGGA TAAACCTGTC ATAGAGTCTC TCAAATGTA
CAGCACGAAA GTGGAAACAC CTTGGGTTG TGA

(SEQ ID NO 196)

Figure 100

TAAGGATAAG GAAGAAGCCT GAGAAGGTTT CTGACTAGGT TGGCAAGCA TTATATGTA AGAGCAAGCA
TTCTATTCA TTTGTGTTGT TAAGAGTAGC GCGGTGAGGA CGAGACATAT AGTTTGAT CAAGTATGTT
ATTGTAAGA ATAATCATG GTAACAAGTA TATTCAAGC ATAATAATAG ACGTAAAGA GTATTTGTCT
TTTAGGTGAA GTGCTTGCAT GGATCTATAG AAATTACA

(SEQ ID NO 197)

Figure 101

TAAGGATAAG GAAACCTGTG AATCTTTTC CCTTCTTTG TTCAGTTTG AGAGGTTCAT CTCTCAAAC
GTGTTCTTG AAAACTAGAT AAGAAAAGTT AGTGTAAAAA GACGAAAGAGA AACCGTAGGT TTTTCTTCAA
CCAAACCGA GAATCAAACC GAGAAAGAAT CTTTCCGTT TCATAAGCGA TCGCACGTT ATGAAAACAC
AACAAACACCT TCGTAAGAAG GATGA

(SEQ ID NO 213)

Figure 102

TAAGGATAAG GAAACCTGTG AATCTTTTC CCTTCTTTG TTCAGTTTG AGAGGTCAAT GACGGCTCAT
CTGAGTACCA GGTGACACGT TTTTGAGGTG TCTCTTCGTA TGAGGGCCT ATAGCTCAGC TGGTTAGAGC
GCACGGCTGA TAAGCGTAG GTCGGTGGTT CGAGTCCACT TAGGCCACT TTTTGAATA AACCTTTCTT
TTTATATGT TAATAAGGG CCTTAGCTCA GCTGGGAGAG CGCCTGCTT GCACCCAGGA GGTCAAGCGGT
TCGATCCGGC TAGGCTCCAC CAAAGATAGT TTGTTCTTTG AAAACTAGAT AAGAAAGTT AGTGTAAAAA
GACGAAGAGA AACCGTAGGT TTTCTCAA CCAAAACCGA GAATCAAACC GAGAAAGAAT CTTTCCGTT
TCATAAGCGA TCGCACCGTT ATGAAAACAC AACAACACCT TCGTAAGAAG GATGA

(SEQ ID NO 214)

Figure 103

TAAGGATAAG GAAACCTGTG AATCTTTTC CCTTCTTTTG TTCAGTTTG AGAGGGCAAT GACGGCTCATA
CTGAGTACCA CGTGACACGT TTTTGAGGTG TCTCTCGTA TGAGGGCCT ATAGCTCAGC TGGTTAGAGC
GCACGGCCTGA TAAGGGTGTG GTCGGGTGGTT CGAGTCCACT TAGGCCCACT TTTTGAATA AACCTTTCTT
TTTTATATGT TAATAAGGG CCTTAGCTCA GCTGGGAGAG CGCCTGCTTT GCACGCAGGA GGTCAAGGGT
TCGATCCCGC TAGGCTCCAC CAAAGATACT TTGTTCTTTG AAAACTAGAT AAGAAAAGTT AGTGTAAAAA
GACGAAGAGA AACCGTAGGT TTTCTCAA CCAAACCGA GAAAGAATCT TTCCGTTTC ATAAGGGATC
GCACGGTTAT GAAAACACAA CAACACCTTC GTAAGAAGGA TGA

(SEQ ID NO 215)

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(54) **Simultaneous detection, identification and differentiation of Eubacterial taxa using a hybridization assay**

(57) The present invention relates to a method for detection and identification of at least one micro-organism, or for the simultaneous detection of several micro-organisms in a sample, comprising the steps of:

- (i) if need be releasing, isolating or concentrating the polynucleic acids present in the sample;
- (ii) if need be amplifying the 16S-23S rRNA spacer region, or a part of it, with at least one suitable primer pair;
- (iii) hybridizing the polynucleic acids of step (i) or (ii) with at least one and preferably more than one of the spacer probes as mentioned in table Ia or equivalents of thereof, under the appropriate hybridization

and wash conditions, and/or with a taxon-specific probe derived from any of the spacer sequences as represented in figs. 1-103 under the same hybridization and wash conditions;

- (iv) detecting the hybrids formed in step (iii) with each of the probes used under appropriate hybridization and wash conditions;
- (v) identification of the micro-organism(s) present in the sample from the differential hybridization signals obtained in step (iv).

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EUROPEAN SEARCH REPORT

Application Number
EP 01 20 0037

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